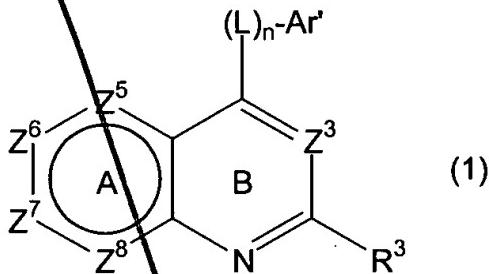


**In the Claims:**

Please amend the claims as follows:

**Please replace the presently pending claims with the following claims:**

1. (Amended) A method to inhibit p38 $\alpha$  activity, which method comprises contacting said p38 $\alpha$  with a compound of the formula:



or the pharmaceutically acceptable salts thereof

wherein R<sup>3</sup> comprises a substituted or unsubstituted aromatic moiety, wherein said aromatic moiety is a monocyclic or fused bicyclic moiety containing 5-12 ring member atoms, optionally comprising one or more heteroatoms selected from O, S and N;

each Z is CR<sup>2</sup> or N, wherein no more than two Z positions in ring A are N, and wherein two adjacent Z positions in ring A cannot be N;

each R<sup>2</sup> is either

(i) independently selected from the group consisting of H, alkyl, alkenyl, alkynyl, acyl, wherein each of alkyl, alkenyl, alkynyl and acyl may optionally contain 1-2 O, S or N, aryl, and arylalkyl, each of said aryl and arylalkyl optionally containing 1 or more O, S or N and wherein in each of the foregoing other than H may be unsubstituted or substituted with 1-3 substituents selected independently from the group consisting of alkyl, alkenyl, alkynyl, aryl, alkylaryl, aroyl, N-aryl, NH-alkylaryl, NH-aroyl, halo, OR, NR<sub>2</sub>, SR, -SOR, -SO<sub>2</sub>R, -OCOR, -NRCOR, -NRCONR<sub>2</sub>, -NRCOOR, -NRSOR, -NRSO<sub>2</sub>R, -OCONR<sub>2</sub>, RCO, -COOR, -SO<sub>3</sub>R, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or alkyl (1-4C), and wherein any aryl or aroyl groups on said substituents may be further substituted by alkyl, alkenyl, alkynyl, halo, OR, NR<sub>2</sub>, SR, -SOR, -SO<sub>2</sub>R, -OCOR, -NRCOR, -NRCONR<sub>2</sub>, -NRCOOR, -NRSOR, -NRSO<sub>2</sub>R, -OCONR<sub>2</sub>, RCO, -COOR, -SO<sub>3</sub>R, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or alkyl (1-4C), or

~~(ii) independently selected from the group consisting of halo, OR, NR<sub>2</sub>, SR, -SOR, -SO<sub>2</sub>R, -OCOR, -NRCOR, -NRCONR<sub>2</sub>, -NRCOOR, NRSOR, NRSO<sub>2</sub>R, -OCONR<sub>2</sub>, RCO, -COOR, -SO<sub>3</sub>R, NRSOR, NRSO<sub>2</sub>R, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or alkyl (1-4C);~~

*A3*  
*B'*  
*cont*  
*D*  
*E*  
*F*  
*G*  
*H*  
*I*  
*J*  
*K*  
*L*  
*M*  
*N*  
*O*  
*P*  
*Q*  
*R*  
*S*  
*T*  
*U*  
*V*  
*W*  
*X*  
*Y*  
*Z*

wherein L is a divalent moiety that provides a distance of 2-8Å between ring B and Ar'; n is 0 or 1; and

Ar' is a cyclic aliphatic, cyclic heteroaliphatic or a monocyclic or polycyclic aromatic moiety any of the foregoing optionally substituted with 1-3 substituents, wherein two of said substituents may form a 5-7 member cyclic optionally heterocyclic aliphatic ring and wherein Ar' and any said substituents thereon forming a cyclic aliphatic ring, may optionally contain one or more ring atoms selected from O, S and N.

Please cancel claims 2-7.

8. (Amended) The method of claim 1 wherein any substituents on the aromatic or heteroaromatic moiety of R<sup>3</sup> are independently selected from the group consisting of halo, OR, NR<sub>2</sub>, SR, -SOR, -SO<sub>2</sub>R, -OCOR, -NRCOR, -NRCONR<sub>2</sub>, -NRCOOR, -NRSOR, -NRSO<sub>2</sub>R, -OCONR<sub>2</sub>, RCO, -COOR, -SO<sub>3</sub>R, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or alkyl (1-4C) and alkyl (1-6C).

9. The method of claim 1 wherein said substituents on substituted Ar' are independently selected from the group consisting of optionally substituted alkyl, alkenyl, alkynyl, aryl, alkylaryl, aroyl, N-aryl, NH-alkylaryl, NH-aroyl, halo, OR, NR<sub>2</sub>, SR, -SOR, -SO<sub>2</sub>R, -OCOR, -NRCOR, -NRCONR<sub>2</sub>, -NRCOOR, -NRSOR, -NRSO<sub>2</sub>R, -OCONR<sub>2</sub>, RCO, -COOR, -SO<sub>3</sub>R, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or alkyl (1-4C),

and wherein any aryl or aroyl groups on said substituents may be further substituted by alkyl, alkenyl, alkynyl, halo, OR, NR<sub>2</sub>, SR, -SOR, -SO<sub>2</sub>R, -OCOR, -NRCOR, -NRCONR<sub>2</sub>, -NRCOOR, -NRSOR, -NRSO<sub>2</sub>R, -OCONR<sub>2</sub>, RCO, -COOR, -SO<sub>3</sub>R, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or alkyl (1-4C).

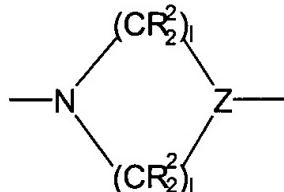
A4  
10. (Amended) The method of claim 9 wherein Ar' is phenyl, 2-, 3-, or 4-pyridyl, 2- or 4-pyrimidyl, indolyl, isoquinolyl, quinolyl, benzimidazolyl, benzotriazolyl, benzothiazolyl, benzofuranyl, pyridyl, thienyl, furyl, pyrrolyl, thiazolyl, oxazolyl, or imidazolyl, all of which may optionally be substituted.

Please cancel claims 11 and 12.

A5  
13. (Amended) The method of claim 1 wherein said optional substituents on R<sup>2</sup> are independently selected from the group consisting of R<sup>4</sup>, halo, OR<sup>4</sup>, NR<sup>4</sup><sub>2</sub>, SR<sup>4</sup>, -OOCR<sup>4</sup>, -NROCR<sup>4</sup>, -COOR<sup>4</sup>, R<sup>4</sup>CO, -CONR<sup>4</sup><sub>2</sub>, -SO<sub>2</sub>NR<sup>4</sup><sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R<sup>4</sup> is independently H, or optionally substituted alkyl (1-6C), or optionally substituted arylalkyl (7-12C) and wherein two R<sup>4</sup> or two substituents on said alkyl or arylalkyl taken together may form a fused aliphatic ring of 5-7 members.

Please cancel claim 14.

A6  
15. (Amended) The method of claim 1 wherein L is S(CR<sup>2</sup><sub>2</sub>)<sub>m</sub>, -NR<sup>1</sup>SO<sub>2</sub>(CR<sup>2</sup><sub>2</sub>)<sub>l</sub>, SO<sub>2</sub>(CR<sup>2</sup><sub>2</sub>)<sub>m</sub>, SO<sub>2</sub>NR<sup>1</sup>(CR<sup>2</sup><sub>2</sub>)<sub>l</sub>, NR<sup>1</sup>(CR<sup>2</sup><sub>2</sub>)<sub>m</sub>, NR<sup>1</sup>CO(CR<sup>2</sup><sub>2</sub>)<sub>l</sub>, O(CR<sup>2</sup><sub>2</sub>)<sub>m</sub>, or OCO(CR<sup>2</sup><sub>2</sub>)<sub>l</sub>, or



wherein Z is N or CH and wherein m is 0-4 and l is 0-3;

R<sup>1</sup> is H, alkyl or arylalkyl where the aryl moiety may be substituted by 1-3 substituents selected independently from the group consisting of alkyl, alkenyl, alkynyl, aryl, alkylaryl, aroyl, N-aryl, NH-alkylaryl, NH-aroyl, halo, OR, NR<sub>2</sub>, SR, -SOR, -SO<sub>2</sub>R, -OCOR, -NRCOR, -NRCONR<sub>2</sub>, -NRCOOR, -NRSOR, -NRSO<sub>2</sub>R, -OCONR<sub>2</sub>, RCO, -COOR, -SO<sub>3</sub>R, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or alkyl (1-4C);

and wherein any aryl or aroyl groups on said substituents may be further substituted by alkyl, alkenyl, alkynyl, halo, OR, NR<sub>2</sub>, SR, -SOR, -SO<sub>2</sub>R, -OCOR, -NRCOR, -NRCONR<sub>2</sub>,

-NRCOOR, -NRSOR, -NRSO<sub>2</sub>R, -OCONR<sub>2</sub>, RCO, -COOR, -SO<sub>3</sub>R, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or alkyl (1-4C); and

R<sup>2</sup> is as defined in claim 1.

16. (Amended) The method of claim 1 wherein the compound of formula (1) is selected from group consisting of

(a) the compounds listed in Table 2 below, wherein Z<sup>5</sup>-Z<sup>8</sup> are CH; Z<sup>3</sup> is N; R<sup>1</sup> in compound No. 11 is 2-propyl; R<sup>1</sup> in compound No. 12 is 4-methoxyphenyl, and R<sup>1</sup> in compound No. 41 is 4-methoxybenzyl; and wherein L, Ar' and R<sup>3</sup> are as shown in Table 2:

A6  
Table 2

Compound No.	L	Ar'	R <sup>3</sup>
1	NH	4-pyridyl	2-chlorophenyl
2	NH	4-pyridyl	2,6-dichlorophenyl
3	NH	4-pyridyl	2-methylphenyl
4	NH	4-pyridyl	2-bromophenyl
5	NH	4-pyridyl	2-fluorophenyl
6	NH	4-pyridyl	2,6-difluorophenyl
7	NH	4-pyridyl	phenyl
8	NH	4-pyridyl	4-fluorophenyl
9	NH	4-pyridyl	4-methoxyphenyl
10	NH	4-pyridyl	3-fluorophenyl
11	NR <sup>1</sup>	4-pyridyl	phenyl
12	NR <sup>1</sup>	4-pyridyl	phenyl
13	NHCH <sub>2</sub>	4-pyridyl	phenyl
14	NHCH <sub>2</sub>	4-pyridyl	4-chlorophenyl
15	NH	3-pyridyl	phenyl
16	NHCH <sub>2</sub>	2-pyridyl	phenyl
17	NHCH <sub>2</sub>	3-pyridyl	phenyl
18	NHCH <sub>2</sub>	2-pyridyl	phenyl
19	NHCH <sub>2</sub> CH <sub>2</sub>	2-pyridyl	phenyl
20	NH	6-pyrimidinyl	phenyl
21	NH	2-pyrimidinyl	phenyl
22	NH	Phenyl	phenyl
23	NHCH <sub>2</sub>	Phenyl	3-chlorophenyl
24	NH	3-hydroxyphenyl	phenyl
25	NH	2-hydroxyphenyl	phenyl

Table 2

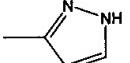
Compound No.	L	Ar'	R <sup>3</sup>
26	NH	4-hydroxyphenyl	phenyl
27	NH	4-indolyl	phenyl
28	NH	5-indolyl	phenyl
29	NH	4-methoxyphenyl	phenyl
30	NH	3-methoxyphenyl	phenyl
31	NH	2-methoxyphenyl	phenyl
32	NH	4-(2-hydroxyethyl)phenyl	phenyl
33	NH	3-cyanophenyl	phenyl
34	NHCH <sub>2</sub>	2,5-difluorophenyl	phenyl
35	NH	4-(2-butyl)phenyl	phenyl
36	NHCH <sub>2</sub>	4-dimethylaminophenyl	phenyl
38	NH	2-pyridyl	phenyl
39	NHCH <sub>2</sub>	3-pyridyl	phenyl
40	NH	4-pyrimidyl	phenyl
41	NR <sup>1</sup>	4-pyridyl	phenyl
42	NH	p-aminomethylphenyl	phenyl
43	NHCH <sub>2</sub>	4-aminophenyl	phenyl
44	NH	4-pyridyl	3-chlorophenyl
45	NH	Phenyl	4-pyridyl
46	NH		phenyl
48	NH	2-benzylamino-3-pyridyl	phenyl
49	NH	2-benzylamino-4-pyridyl	phenyl
50	NH	3-benzyloxyphenyl	phenyl
51	NH	4-pyridyl	3-aminophenyl
52	NH	4-pyridyl	4-pyridyl
53	NH	4-pyridyl	2-naphthyl
54		4-pyridyl	phenyl
55		Phenyl	phenyl
56		2-pyridyl	phenyl
61	NH	4-pyridyl	2-trifluoromethyl phenyl
62	NH	4-aminophenyl	phenyl
64	NH	3-methoxyphenyl	2-fluorophenyl
65	NH	4-methoxyphenyl	2-fluorophenyl

Table 2

Compound No.	L	Ar'	R <sup>3</sup>
66	NH	4-pyrimidinyl	2-fluorophenyl
67	NH	3-amino-4-pyridyl	phenyl
68	NH	4-pyridyl	2-benzylaminophenyl
69	NH	2-benzylaminophenyl	phenyl
70	NH	2-benzylaminophenyl	4-cyanophenyl
71	NH	3'-cyano-2-benzylaminophenyl	phenyl

A6

(b) the compounds listed in Table 3, below, wherein L is NH; Z<sup>3</sup> is N; Z<sup>6</sup> and Z<sup>7</sup> are CH and Z<sup>5</sup>, Z<sup>8</sup>, Ar' and R<sup>3</sup> are as shown in Table 3:

Table 3

Compound No.	Z <sup>5</sup>	Z <sup>8</sup>	Ar'	R <sup>3</sup>
72	CH	N	4-pyridyl	2-fluorophenyl
73	CH	N	4-pyridyl	2-chlorophenyl
74	CH	N	4-pyridyl	phenyl
75	N	N	4-pyridyl	phenyl
76	N	CH	4-pyridyl	phenyl

and

(c) the quinazoline derivatives listed in Table 4 below, wherein L is NH; Ar' is 4-pyridyl; Z<sup>3</sup>, Z<sup>5</sup>, and Z<sup>8</sup> are N; Z<sup>6</sup> or Z<sup>7</sup> are CR<sup>2</sup> as shown and each is otherwise N and wherein R<sup>3</sup> and R<sup>2</sup> are as shown in Table 4:

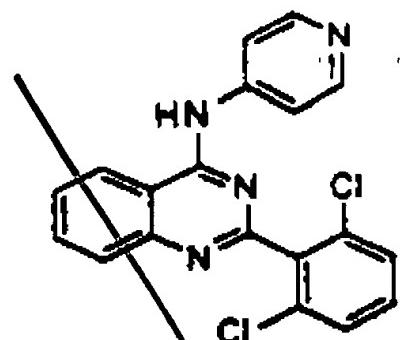
Table 4

Compound No.	R <sup>3</sup>	R <sup>2</sup>
77	2-chlorophenyl	6,7-dimethoxy
78	2-fluorophenyl	6-nitro
79	2-fluorophenyl	6-amino
80	2-fluorophenyl	7-amino
81	2-fluorophenyl	6-(3-methoxybenzylamino)
82	2-fluorophenyl	6-(4-methoxybenzylamino)
83	2-fluorophenyl	6-(2-isobutylamino)
84	2-fluorophenyl	6-(4-methylmercaptophenylamino)
85	2-fluorophenyl	6-(4-methoxybenzoyl amino)
86	4-fluorophenyl	7-amino
87	4-fluorophenyl	7-(3-methoxybenzylamino)

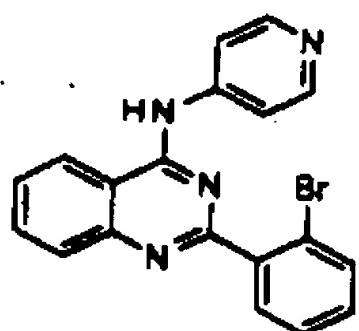
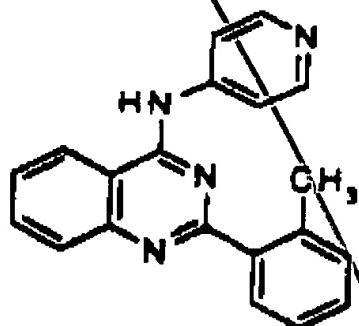
A 6  
 17. (Amended) The method of claim 1 wherein the compound of formula (1) is selected from the group consisting of the following compounds:

B2  
cont

09972582 100501

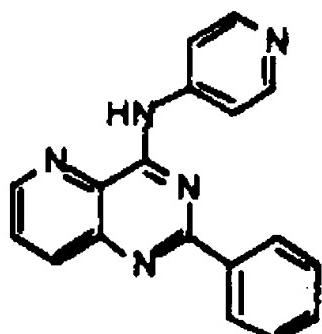
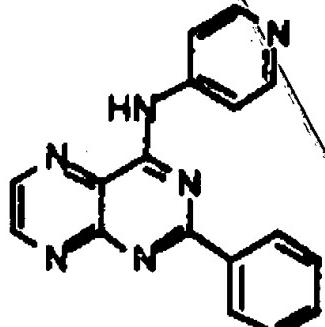
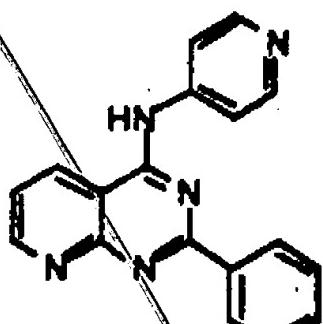
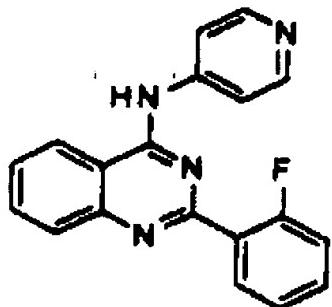


A6



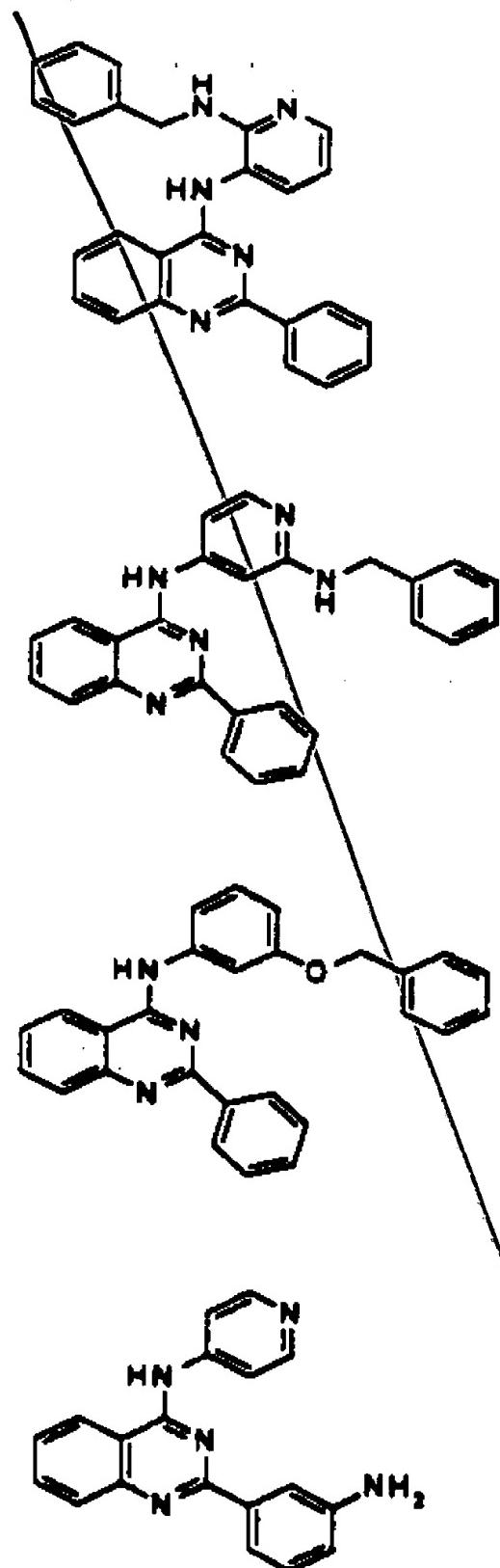
*B<sup>2</sup>*  
*cont*

*A<sup>6</sup>*



B2  
cont

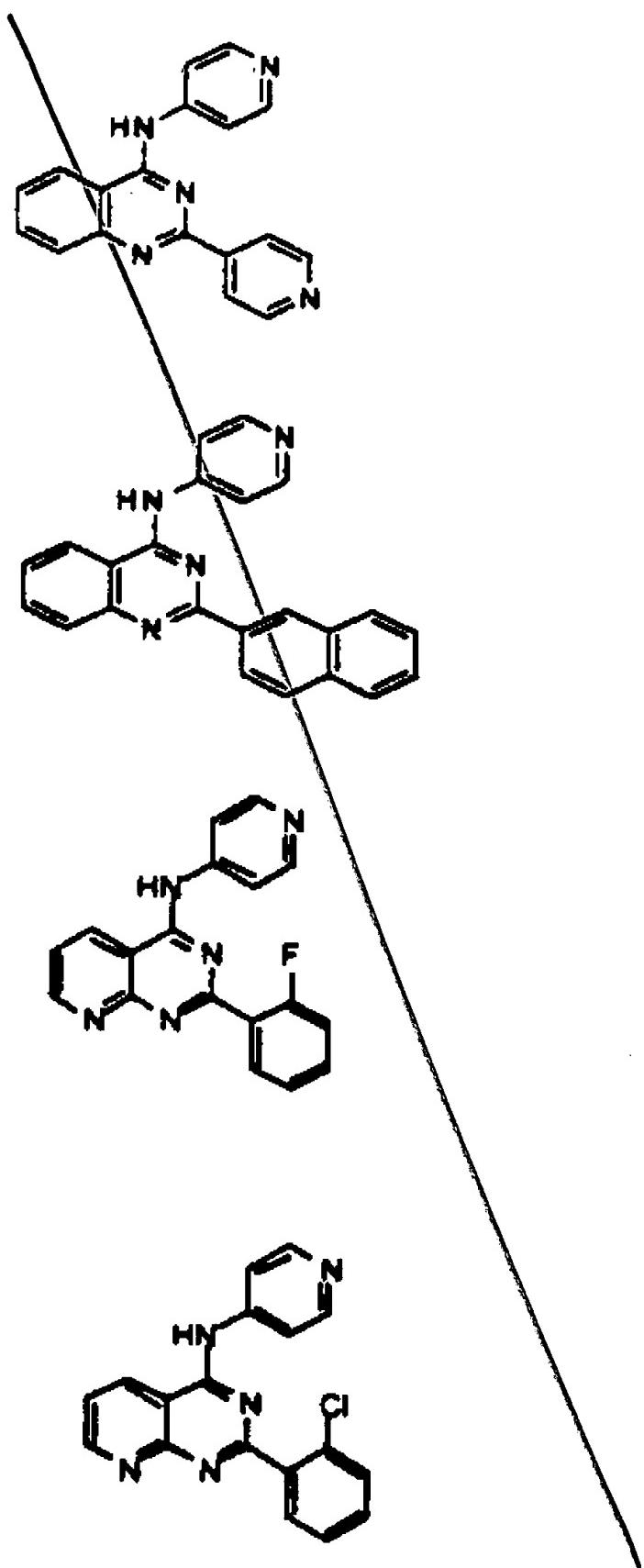
A6



B2  
cont

09972582-100604

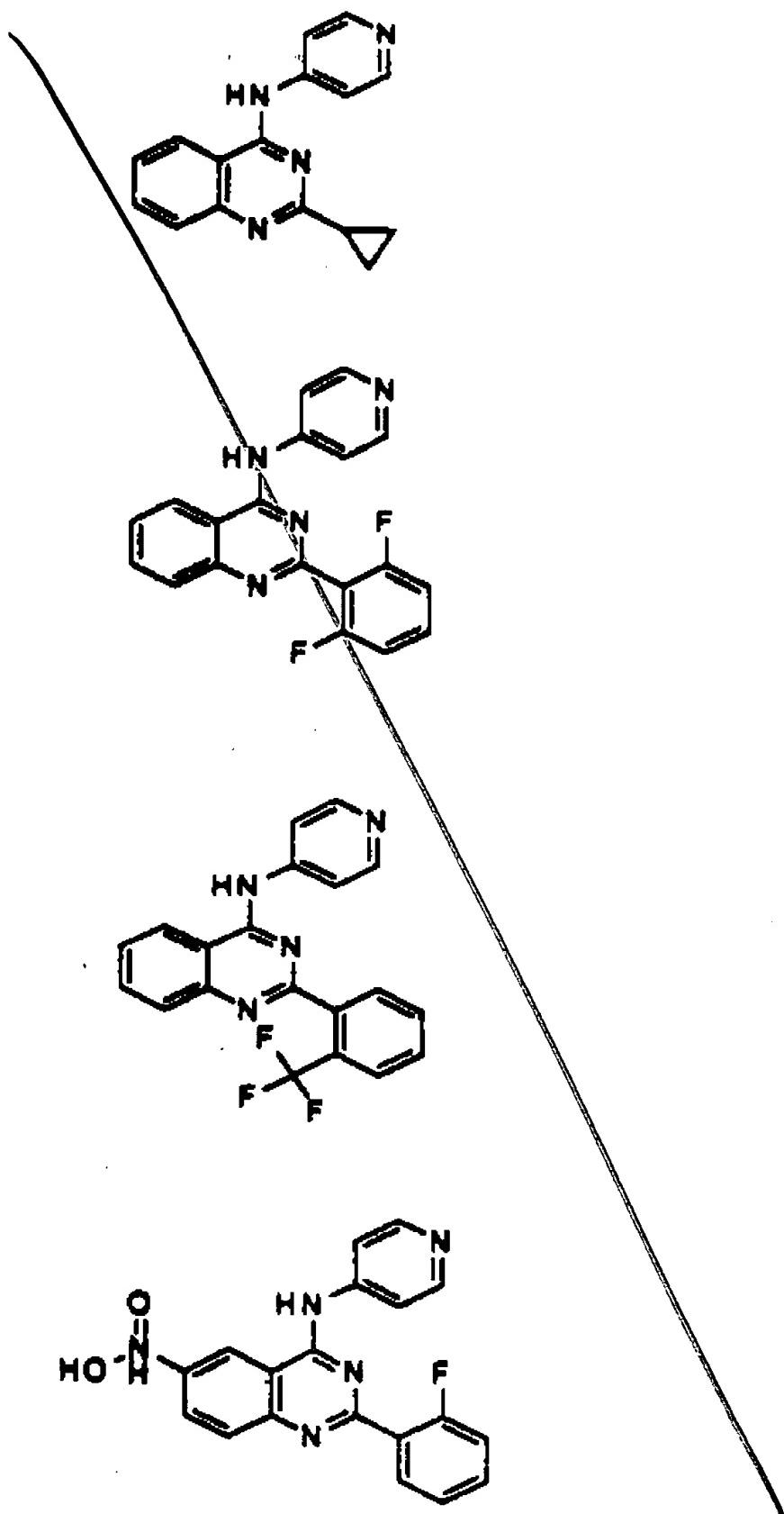
A6



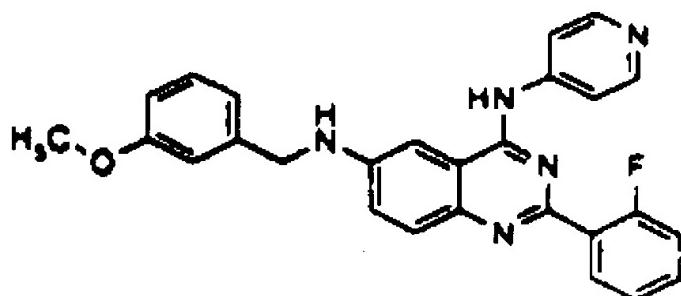
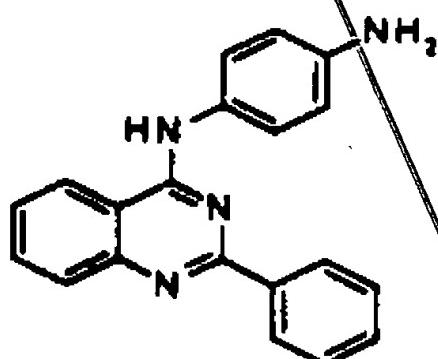
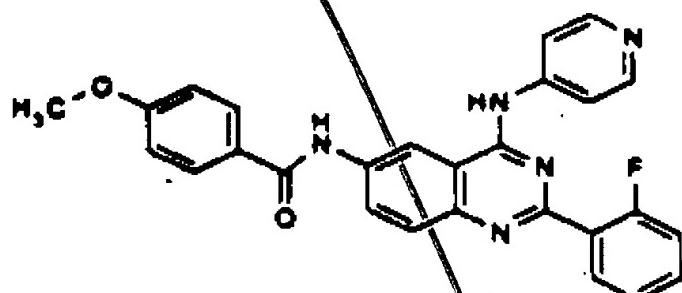
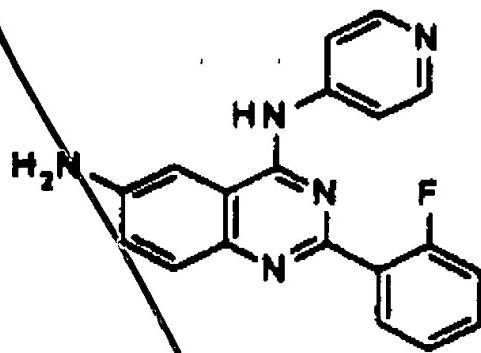
*B<sup>2</sup>*  
cont

09972582-1100501

A6

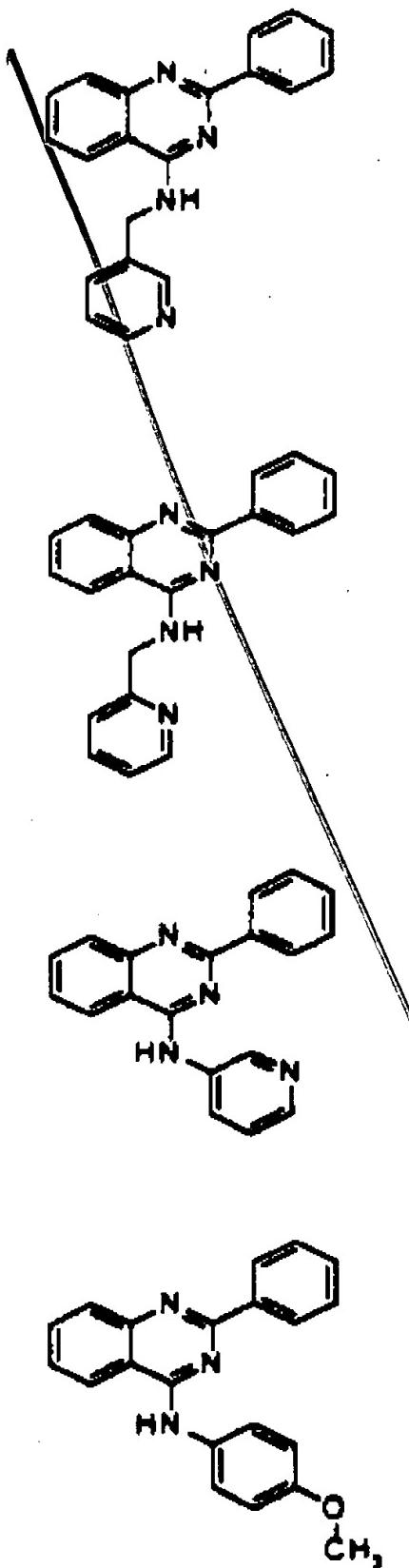


B2  
cont



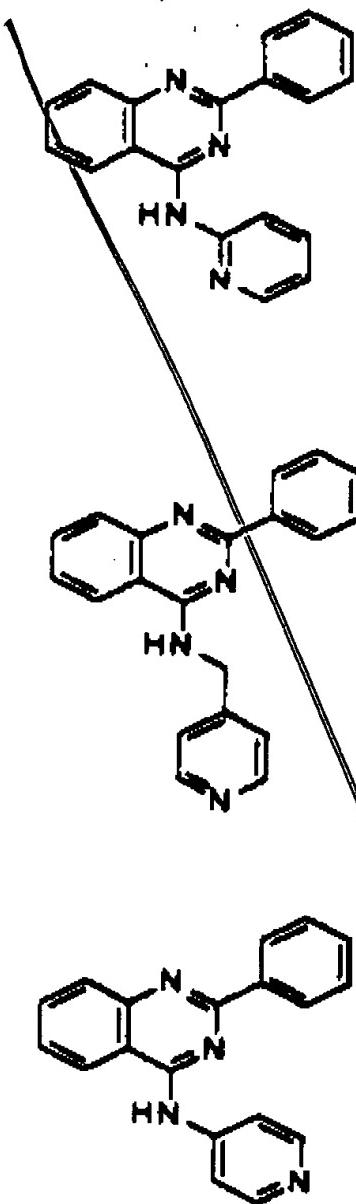
B2  
Cont

A6



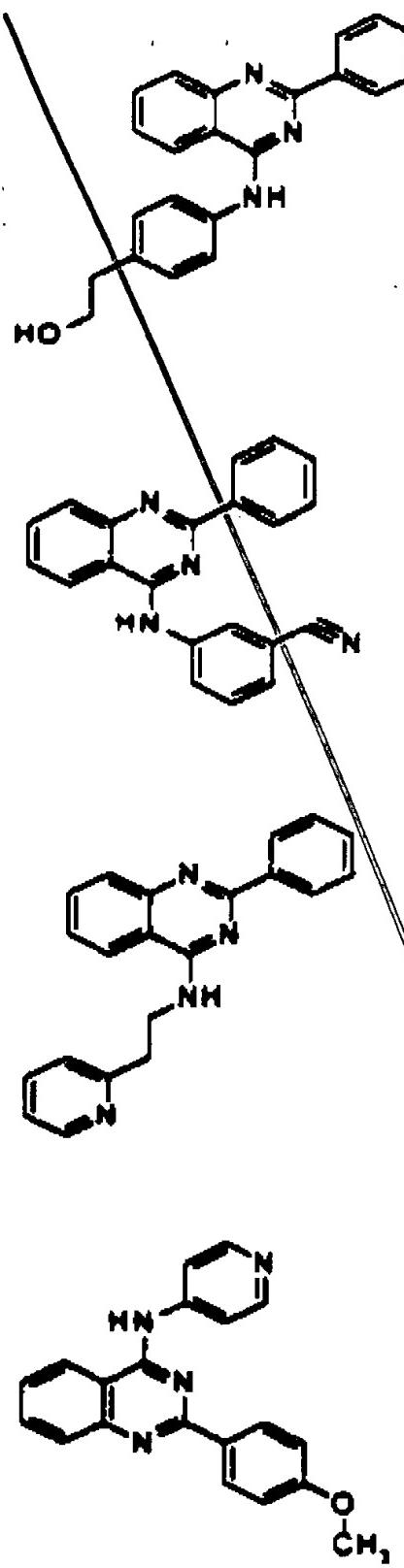
*B2  
cont*

*A6*



B2  
cont

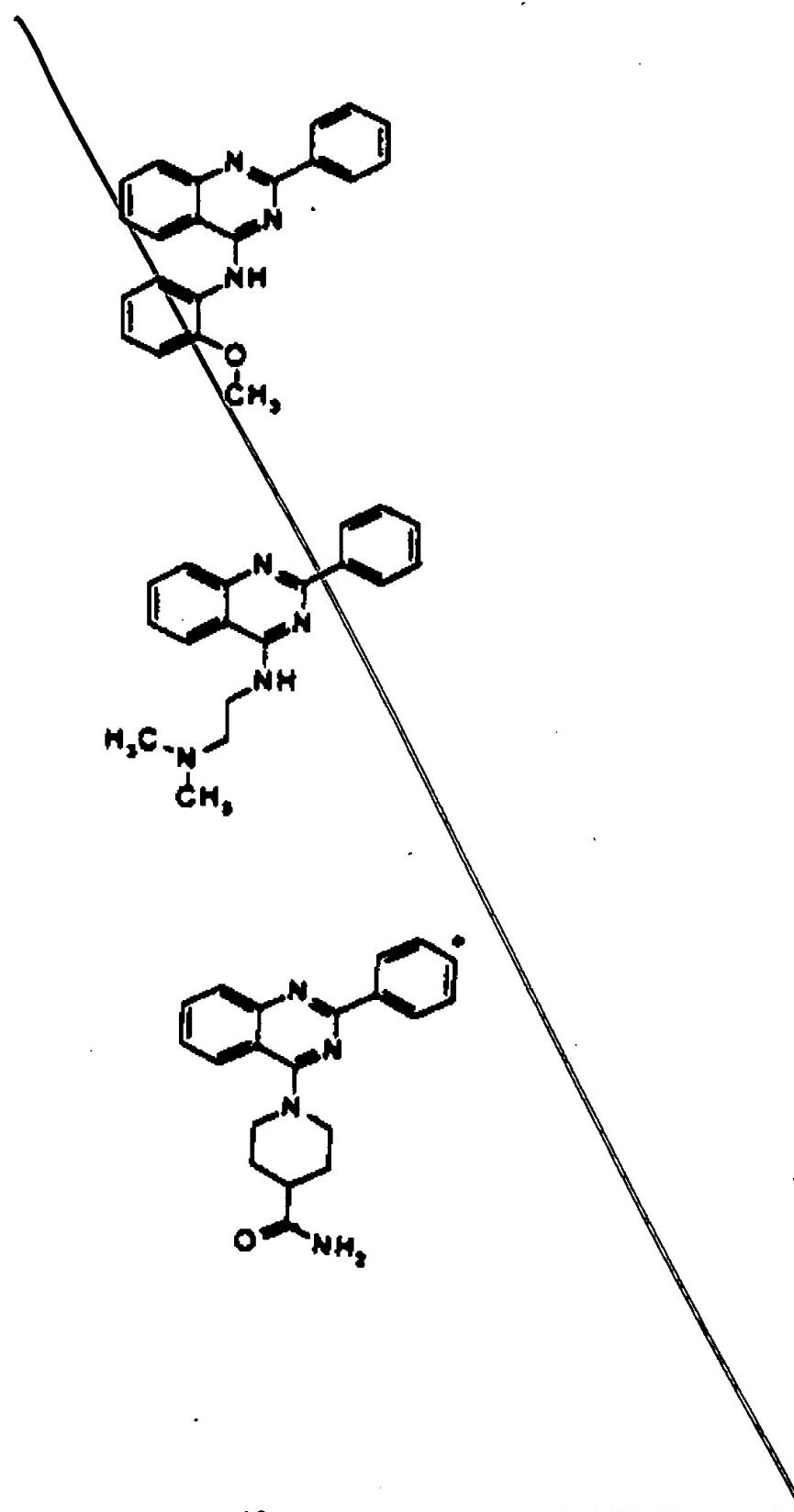
A6



B2  
cont

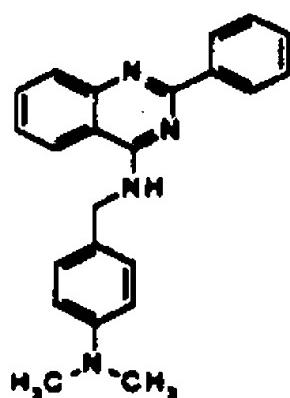
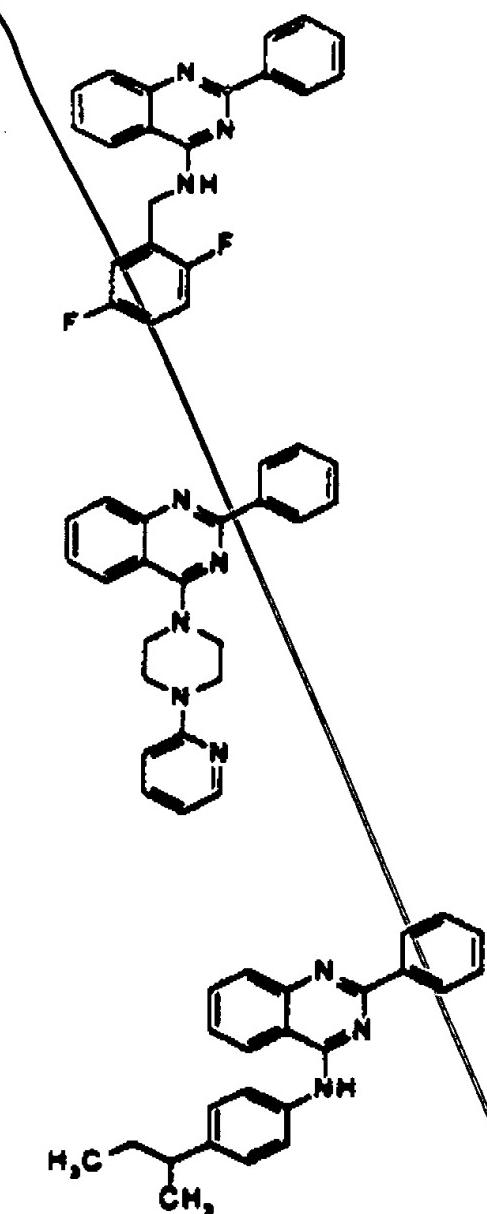
A6

05972532 - 100501



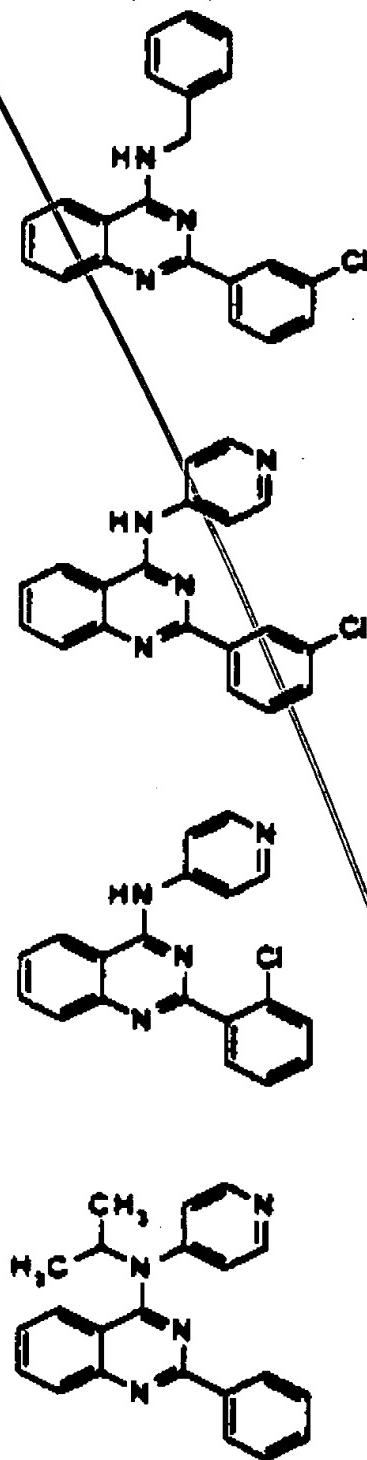
B<sup>2</sup>  
cont

A<sup>b</sup>



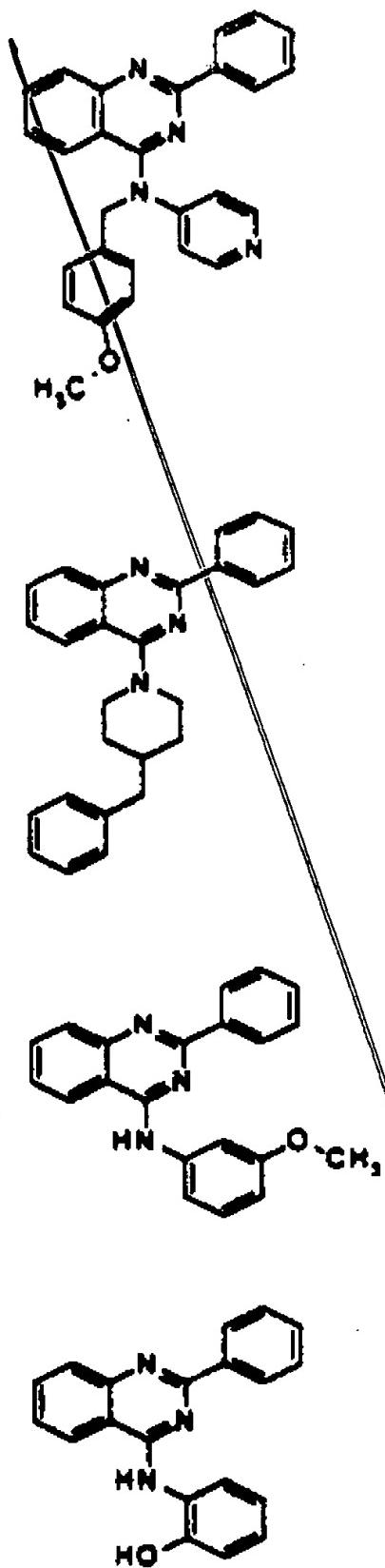
B2  
cont

A6



B<sup>2</sup>  
cont

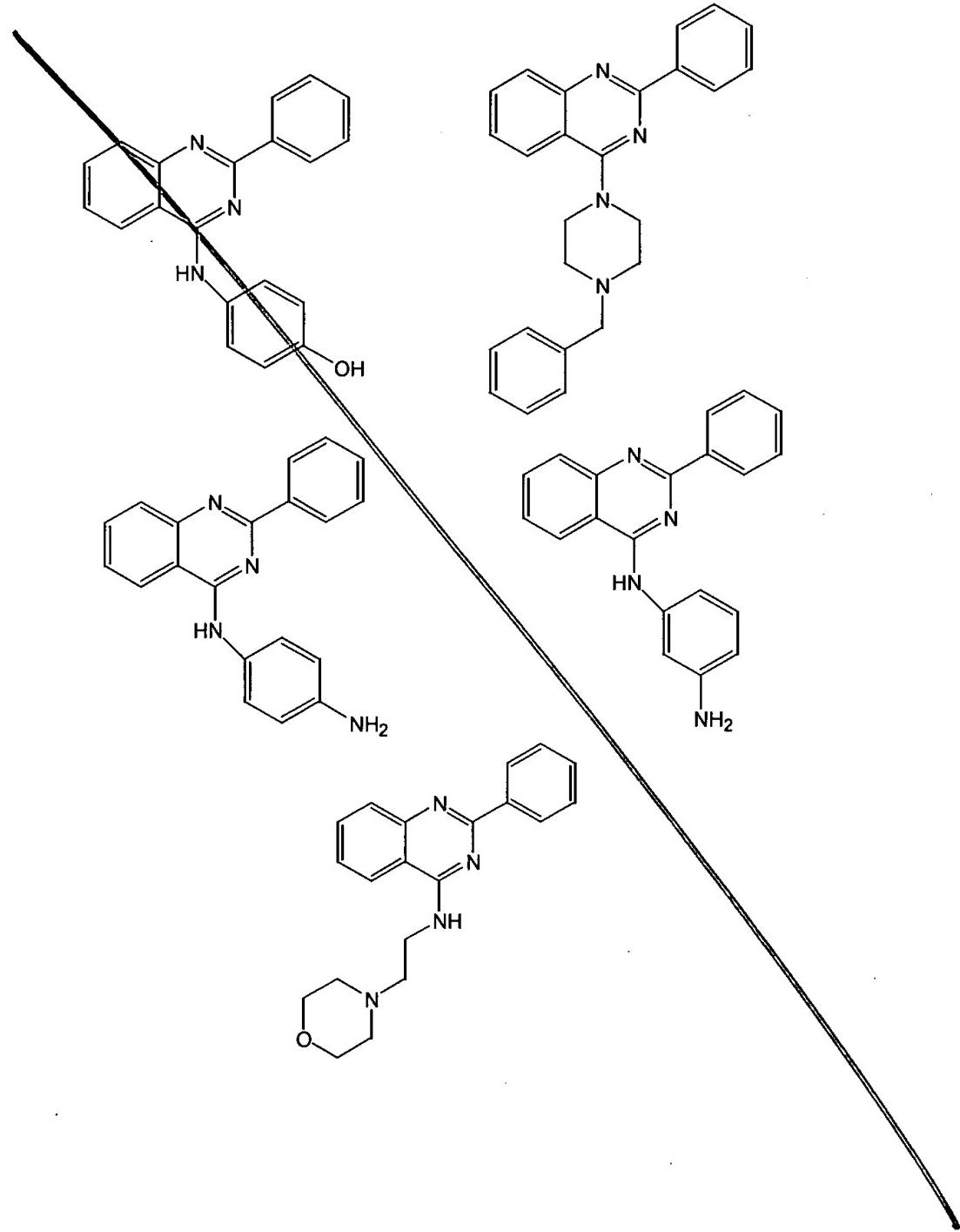
A6



TO50024296

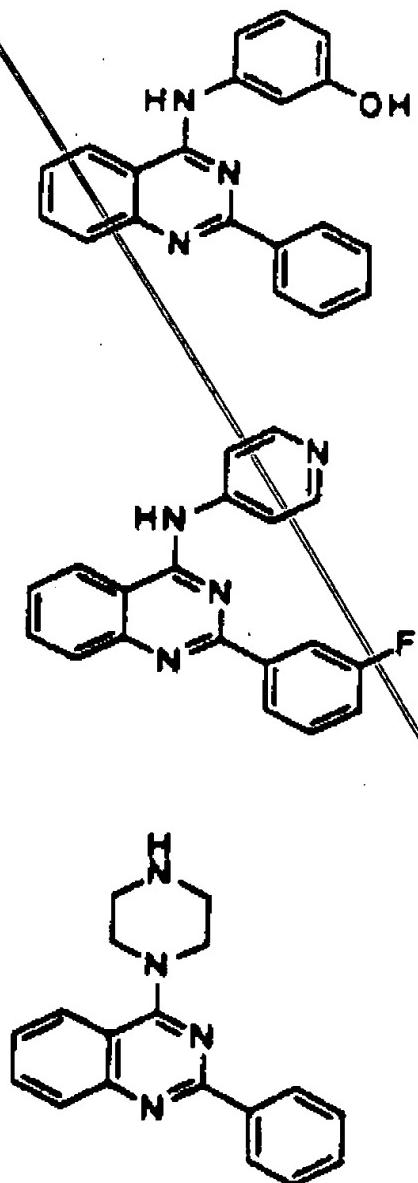
B2  
cont

A6

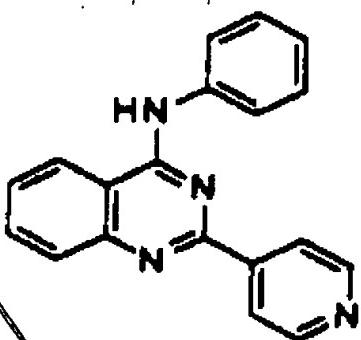


B2  
Bmt

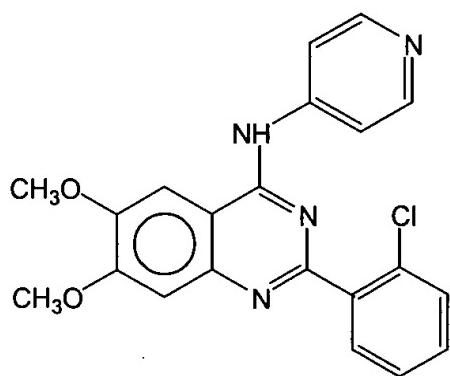
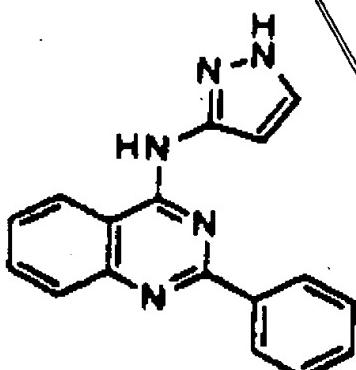
A6



B<sup>2</sup>  
cont



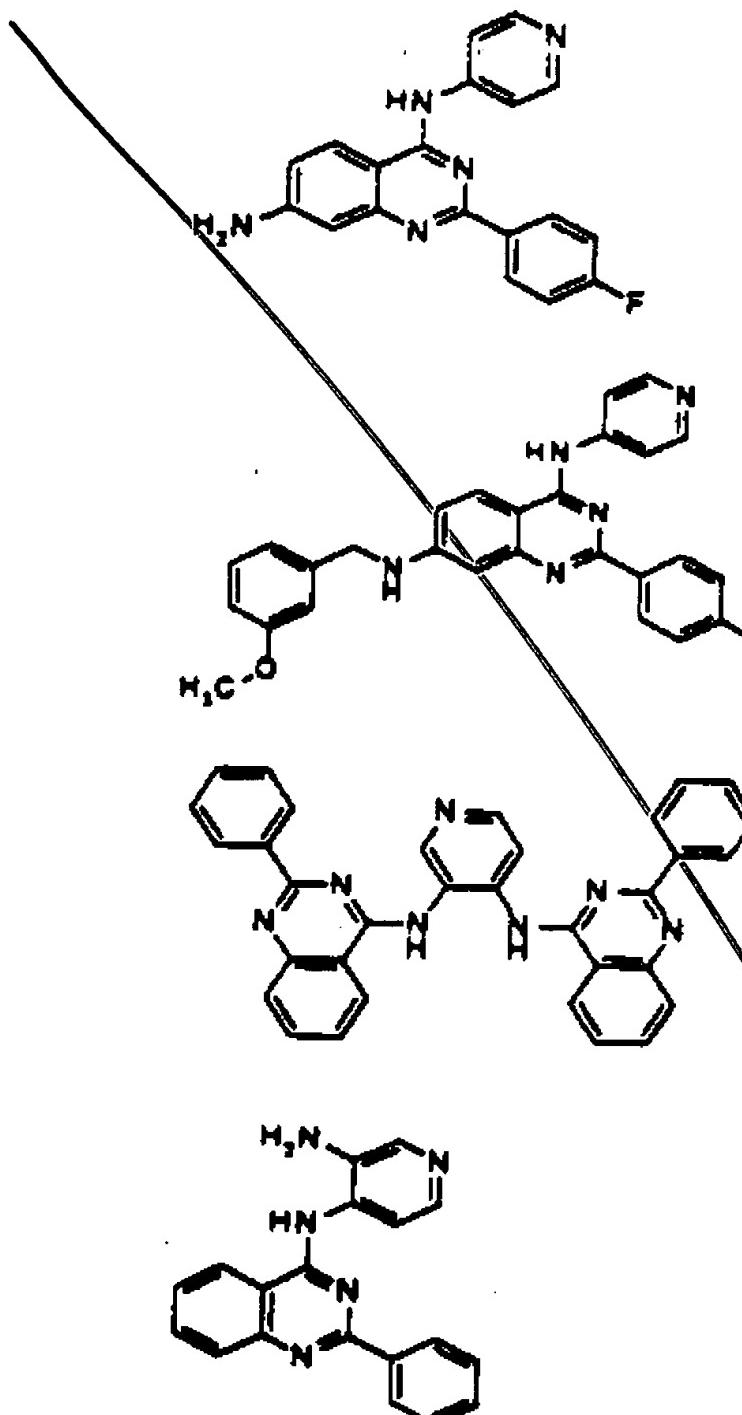
A<sup>b</sup>



B2  
cont

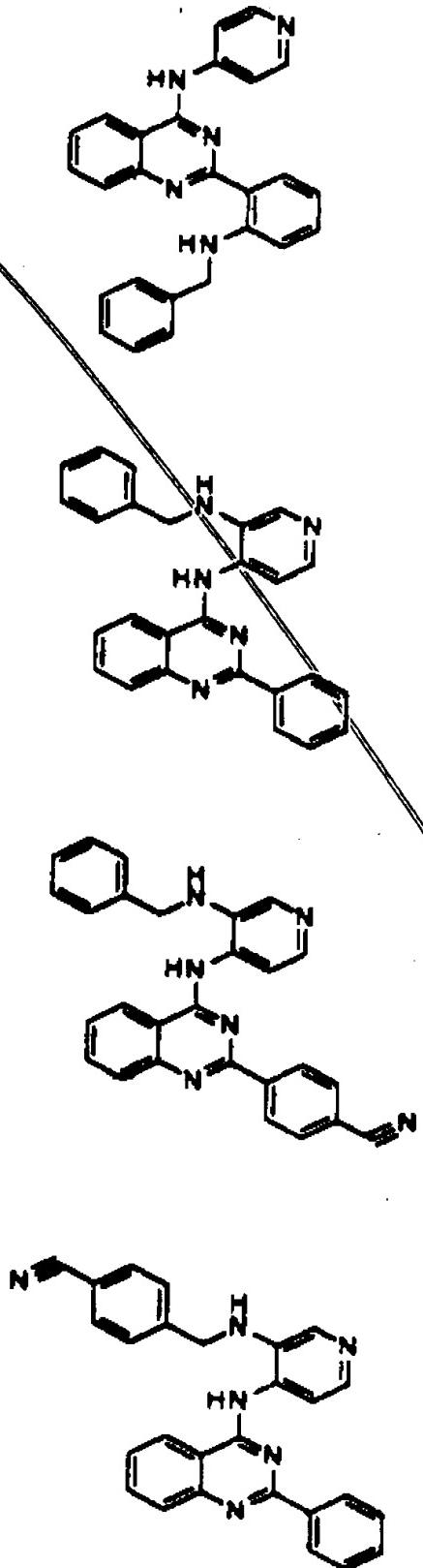
TO500T-22527650

A6



B2  
cont

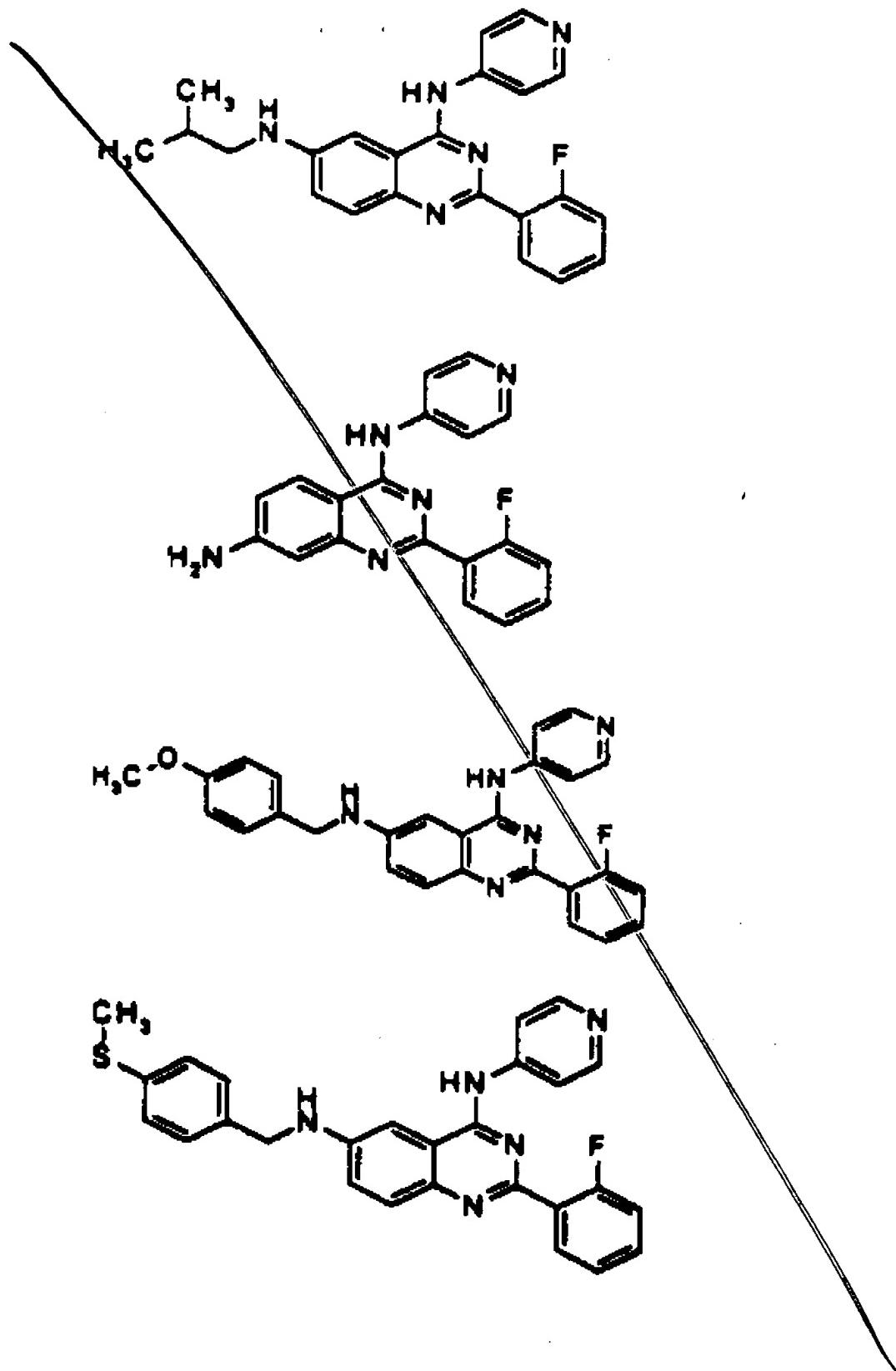
A6



B2  
cont

000072582 = 10001

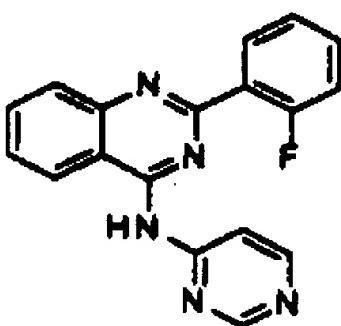
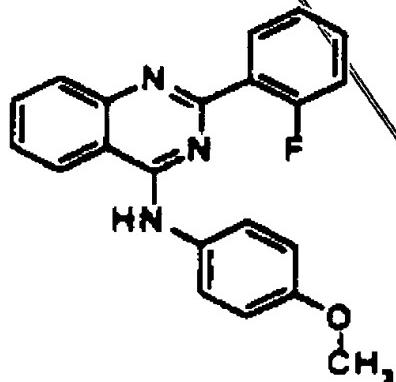
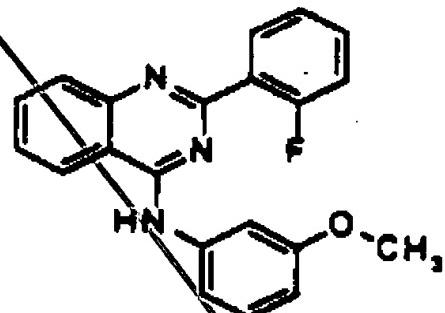
A6



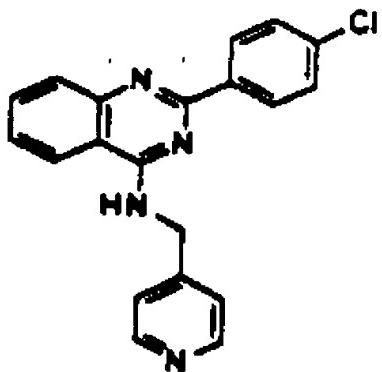
B2  
cont

09972582 = 100501

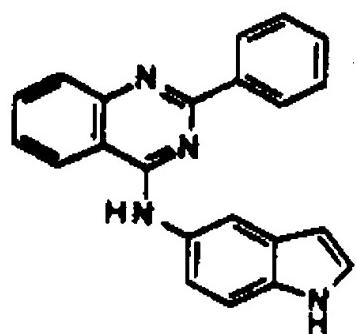
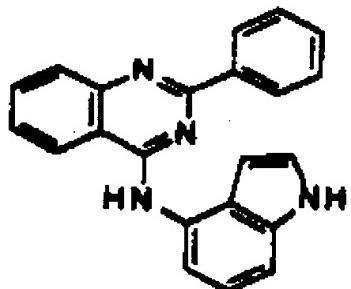
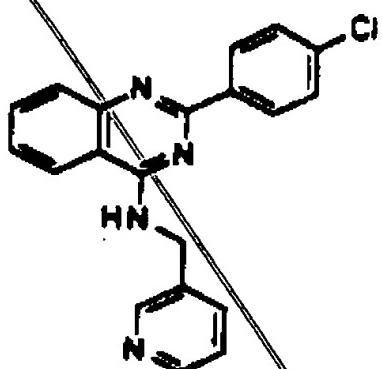
A6



B<sup>2</sup>  
cont



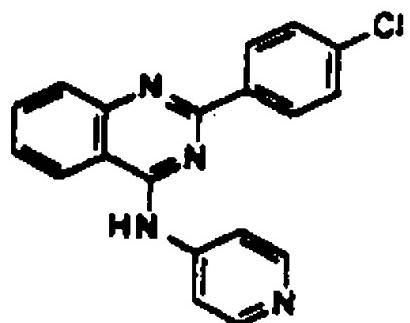
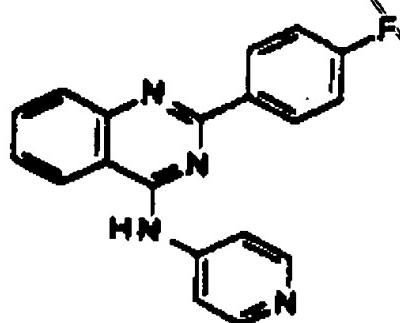
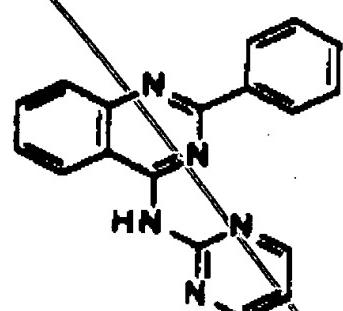
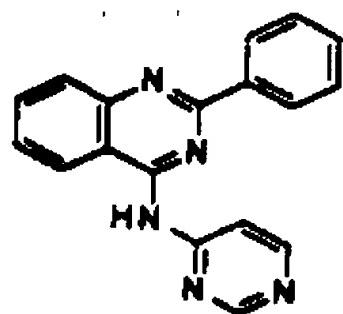
A6



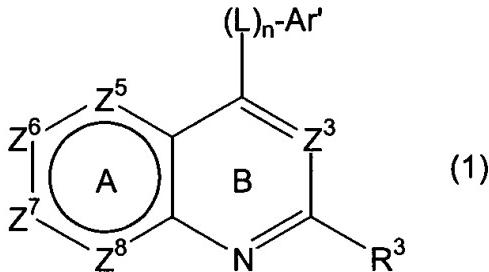
B2  
cont

0099725822 - 4000504

A6



18. (Amended) A pharmaceutical composition for treating conditions characterized by enhanced p38 $\alpha$  kinase activity which composition comprises an amount of a compound of the formula



*A6* or the pharmaceutically acceptable salts thereof

wherein R<sup>3</sup>;

each Z;

each R<sup>2</sup>;

L;

n; and

Ar' are as defined in claim 1 which is effective to inhibit p38 $\alpha$  kinase activity in admixture with at least one pharmaceutically acceptable excipient appropriate for administering to a subject exhibiting enhanced p38 $\alpha$  kinase activity.

19. The composition of claim 18 which further contains an additional therapeutic agent.

20. The composition of claim 19 wherein said additional therapeutic agent is a corticosteroid, a monoclonal antibody, or an inhibitor of cell division.

Please cancel claims 21-22.

Please add the following claims:

23. (New) The method of claim 1 wherein

*A7* L is -R<sup>1</sup>N(CH<sub>2</sub>)<sub>n</sub>- wherein R<sup>1</sup> is H or is alkyl (1-6C) or arylalkyl optionally substituted on the aryl group with 1-3 substituents independently selected from alkyl (1-6C), halo, OR, NR<sub>2</sub>,

SR, -OOCR, -NROCR, RCO, -COOR, -CONR<sub>2</sub>, -SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or lower alkyl (1-4C) and n is 0, 1 or 2; and

(a) Ar' is phenyl, substituted with at least one group selected from the group consisting of optionally substituted alkyl (1-6C), halo, OR, NR<sub>2</sub>, SR, -OOCR, -NROCR, RCO, -COOR, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or lower alkyl (1-4C), or pyridyl, indolyl, or pyrimidyl, each optionally substituted with at least one group selected from the group consisting of optionally substituted alkyl (1-6C), halo, OR, NR<sub>2</sub>, SR, -OOCR, -NROCR, RCO, -COOR, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or lower alkyl (1-4C); and

R<sup>3</sup> is phenyl optionally substituted with 1-3 substituents which substituents are selected from the group consisting of alkyl (1-6C), halo, OR, NR<sub>2</sub>, SR, -OOCR, -NROCR, RCO, -COOR, -CONR<sub>2</sub>, -SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or lower alkyl (1-4C); or

(b) Ar' is phenyl, pyridyl, indolyl, or pyrimidyl, each optionally substituted with a group selected from the group consisting of optionally substituted alkyl (1-6C), halo, OR, NR<sub>2</sub>, SR, -OOCR, -NROCR, RCO, -COOR, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or lower alkyl (1-4C); and

R<sup>3</sup> is phenyl substituted with 1-3 substituents which substituents are selected from the group consisting of alkyl (1-6C), halo, SR, -OOCR, -NROCR, RCO, -COOR, -CONR<sub>2</sub>, -SO<sub>2</sub>NR<sub>2</sub>, CN, and CF<sub>3</sub>, wherein each R is independently H or lower alkyl (1-4C); or

(c) Ar' is phenyl substituted with a group selected from the group consisting of optionally substituted NR<sub>2</sub>, SR, -NROCR, RCO, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, and CF<sub>3</sub>, wherein each R is independently H or lower alkyl (1-4C); or pyridyl substituted with a group selected from the group consisting of optionally substituted alkyl (1-6C), halo, OR, NR<sub>2</sub>, SR, -OOCR, -NROCR, RCO, -COOR, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or lower alkyl (1-4C); or indolyl or pyrimidyl, each optionally substituted with a group selected from the group consisting of optionally substituted alkyl (1-6C), halo, OR, NR<sub>2</sub>, SR, -OOCR, -NROCR, RCO, -COOR, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or lower alkyl (1-4C); and

R<sup>3</sup> is phenyl optionally substituted with 1-3 substituents which substituents are selected from the group consisting of alkyl (1-6C), halo, OR, NR<sub>2</sub>, SR, -OOCR, -NROCR, RCO, -COOR,

-CONR<sub>2</sub>, -SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or lower alkyl (1-4C); or

(d) Ar' is phenyl, pyridyl, indolyl, or pyrimidyl, each optionally substituted with a group selected from the group consisting of optionally substituted alkyl (1-6C), halo, OR, NR<sub>2</sub>, SR, -OOCR, -NROCR, RCO, -COOR, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or lower alkyl (1-4C); and

R<sup>3</sup> is phenyl substituted with 1-3 substituents which substituents are selected from the group consisting of alkyl (1-6C), halo, OR, SR, -OOCR, -NROCR, RCO, -COOR, -CONR<sub>2</sub>, -SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or lower alkyl (1-4C).

A7  
24. (New) The method of claim 1 wherein the compound of formula 1 is selected from the group consisting of

2-phenyl-4-(4-pyridylamino)-quinazoline;  
2-(2-bromophenyl)-4-(4-pyridylamino)-quinazoline;  
2-(2-chlorophenyl)-4-(4-pyridylamino)-quinazoline;  
2-(2-fluorophenyl)-4-(4-pyridylamino)-quinazoline;  
2-(2-methylphenyl)-4-(4-pyridylamino)-quinazoline;  
2-(4-fluorophenyl)-4-(4-pyridylamino)-quinazoline;  
2-(3-methoxyanilyl)-4-(4-pyridylamino)-quinazoline;  
2-(2,6-dichlorophenyl)-4-(4-pyridylamino)-quinazoline;  
2-(2,6-dibromophenyl)-4-(4-pyridylamino)-quinazoline;  
2-(2,6-difluorophenyl)-4-(4-pyridylamino)-quinazoline;  
2-(2-fluorophenyl)-4-(4-pyridylamino)-6, 7-dimethoxyquinazoline;  
2-(4-fluorophenyl)-4-(4-pyridylamino)-6, 7-dimethoxyquinazoline;  
2-(2-fluorophenyl)-4-(4-pyridylamino)-6-nitroquinazoline;  
2-(2-fluorophenyl)-4-(4-pyridylamino)-6-aminoquinazoline;  
2-(2-fluorophenyl)-4-(4-pyridylamino)-7-aminoquinazoline;  
2-(2-fluorophenyl)-4-(4-pyridylamino)-6-(3-methoxybenzylamino)-quinazoline;  
2-(2-fluorophenyl)-4-(4-pyridylamino)-6-(4-methoxybenzylamino)-quinazoline;  
2-(2-fluorophenyl)-4-(4-pyridylamino)-6-(2-isobutylamino)-quinazoline; and  
2-(2-fluorophenyl)-4-(4-pyridylamino)-6-(4-methylmercaptophenylamino)-quinazoline.

A1

25. (New) The composition of claim 18 wherein any substituents on the aromatic or heteroaromatic moiety of R<sup>3</sup> are independently selected from the group consisting of alkyl (1-6C), halo, OR, NR<sub>2</sub>, SR, -SOR, -SO<sub>2</sub>R, -OCOR, -NRCOR, -NRCONR<sub>2</sub>, -NRCOOR, -NRSOR, -NRSO<sub>2</sub>R, -OCONR<sub>2</sub>, RCO, -COOR, -SO<sub>3</sub>R, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or alkyl (1-4C).

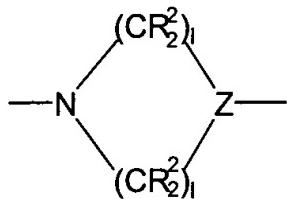
26. (New) The composition of claim 18 wherein said substituents on substituted Ar' are independently selected from the group consisting of optionally substituted alkyl, alkenyl, alkynyl, aryl, alkylaryl, aroyl, N-aryl, NH-alkylaryl, NH-aryloyl, halo, OR, NR<sub>2</sub>, SR, -SOR, -SO<sub>2</sub>R, -OCOR, -NRCOR, -NRCONR<sub>2</sub>, -NRCOOR, -NRSOR, -NRSO<sub>2</sub>R, -OCONR<sub>2</sub>, RCO, -COOR, -SO<sub>3</sub>R, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or alkyl (1-4C),

and wherein any aryl or aroyl groups on said substituents may be further substituted by alkyl, alkenyl, alkynyl, halo, OR, NR<sub>2</sub>, SR, -SOR, -SO<sub>2</sub>R, -OCOR, -NRCOR, -NRCONR<sub>2</sub>, -NRCOOR, -NRSOR, -NRSO<sub>2</sub>R, -OCONR<sub>2</sub>, RCO, -COOR, -SO<sub>3</sub>R, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or alkyl (1-4C).

27. (New) The composition of claim 26 wherein Ar' is phenyl, 2-, 3-, or 4-pyridyl, 2- or 4-pyrimidyl, indolyl, isoquinolyl, quinolyl, benzimidazolyl, benzotriazolyl, benzothiazolyl, benzofuranyl, pyridyl, thienyl, furyl, pyrrolyl, thiazolyl, oxazolyl, or imidazolyl, all of which may optionally be substituted.

28. (New) The composition of claim 18 wherein said optional substituents on R<sup>2</sup> are independently selected from the group consisting of R<sup>4</sup>, halo, OR<sup>4</sup>, NR<sup>4</sup><sub>2</sub>, SR<sup>4</sup>, -OOCR<sup>4</sup>, -NROCR<sup>4</sup>, -COOR<sup>4</sup>, R<sup>4</sup>CO, -CONR<sup>4</sup><sub>2</sub>, -SO<sub>2</sub>NR<sup>4</sup><sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R<sup>4</sup> is independently H, or optionally substituted alkyl (1-6C), or optionally substituted arylalkyl (7-12C) and wherein two R<sup>4</sup> or two substituents on said alkyl or arylalkyl taken together may form a fused aliphatic ring of 5-7 members.

29. (New) The composition of claim 18 wherein L is  $S(CR^2_2)_m$ ,  $-NR^1SO_2(CR^2_2)_l$ ,  $SO_2(CR^2_2)_m$ ,  $SO_2NR^1(CR^2_2)_l$ ,  $NR^1(CR^2_2)_m$ ,  $NR^1CO(CR^2_2)_l$ ,  $O(CR^2_2)_m$ , or  $OCO(CR^2_2)_l$ , or



wherein Z is N or CH and wherein m is 0-4 and l is 0-3;

R<sup>1</sup> is H, alkyl or arylalkyl where the aryl moiety may be substituted by 1-3 substituents selected independently from the group consisting of alkyl, alkenyl, alkynyl, aryl, alkylaryl, aroyl, N-aryl, NH-alkylaryl, NH-aryloyl, halo, OR, NR<sub>2</sub>, SR, -SOR, -SO<sub>2</sub>R, -OCOR, -NRCOR, -NRCONR<sub>2</sub>, -NRCOOR, -NRSOR, -NRSO<sub>2</sub>R, -OCONR<sub>2</sub>, RCO, -COOR, -SO<sub>3</sub>R, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or alkyl (1-4C);

and wherein any aryl or aroyl groups on said substituents may be further substituted by alkyl, alkenyl, alkynyl, halo, OR, NR<sub>2</sub>, SR, -SOR, -SO<sub>2</sub>R, -OCOR, -NRCOR, -NRCONR<sub>2</sub>, -NRCOOR, -NRSOR, -NRSO<sub>2</sub>R, -OCONR<sub>2</sub>, RCO, -COOR, -SO<sub>3</sub>R, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or alkyl (1-4C); and

R<sup>2</sup> is as defined in claim 18.

30. (New) The composition of claim 18 wherein the compound of formula (1) is selected from the group consisting of

(a) the compounds listed in Table 2 below, wherein Z<sup>5</sup>-Z<sup>8</sup> are CH; Z<sup>3</sup> is N; R<sup>1</sup> in compound No. 11 is 2-propyl; R<sup>1</sup> in compound No. 12 is 4-methoxyphenyl, and R<sup>1</sup> in compound No. 41 is 4-methoxybenzyl; and wherein L, Ar' and R<sup>3</sup> are as shown in Table 2:

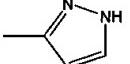
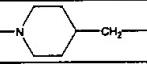
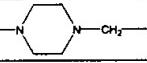
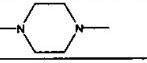
Table 2

Compound No.	L	Ar'	R <sup>3</sup>
1	NH	4-pyridyl	2-chlorophenyl
2	NH	4-pyridyl	2,6-dichlorophenyl
3	NH	4-pyridyl	2-methylphenyl
4	NH	4-pyridyl	2-bromophenyl
5	NH	4-pyridyl	2-fluorophenyl
6	NH	4-pyridyl	2,6-difluorophenyl

Table 2

Compound No.	L	Ar'	R <sup>3</sup>
7	NH	4-pyridyl	phenyl
8	NH	4-pyridyl	4-fluorophenyl
9	NH	4-pyridyl	4-methoxyphenyl
10	NH	4-pyridyl	3-fluorophenyl
11	NR <sup>1</sup>	4-pyridyl	phenyl
12	NR <sup>1</sup>	4-pyridyl	phenyl
13	NHCH <sub>2</sub>	4-pyridyl	phenyl
14	NHCH <sub>2</sub>	4-pyridyl	4-chlorophenyl
15	NH	3-pyridyl	phenyl
16	NHCH <sub>2</sub>	2-pyridyl	phenyl
17	NHCH <sub>2</sub>	3-pyridyl	phenyl
18	NHCH <sub>2</sub>	2-pyridyl	phenyl
19	NHCH <sub>2</sub> CH <sub>2</sub>	2-pyridyl	phenyl
20	NH	6-pyrimidinyl	phenyl
21	NH	2-pyrimidinyl	phenyl
22	NH	Phenyl	phenyl
23	NHCH <sub>2</sub>	Phenyl	3-chlorophenyl
24	NH	3-hydroxyphenyl	phenyl
25	NH	2-hydroxyphenyl	phenyl
26	NH	4-hydroxyphenyl	phenyl
27	NH	4-indolyl	phenyl
28	NH	5-indolyl	phenyl
29	NH	4-methoxyphenyl	phenyl
30	NH	3-methoxyphenyl	phenyl
31	NH	2-methoxyphenyl	phenyl
32	NH	4-(2-hydroxyethyl)phenyl	phenyl
33	NH	3-cyanophenyl	phenyl
34	NHCH <sub>2</sub>	2,5-difluorophenyl	phenyl
35	NH	4-(2-butyl)phenyl	phenyl
36	NHCH <sub>2</sub>	4-dimethylaminophenyl	phenyl
38	NH	2-pyridyl	phenyl
39	NHCH <sub>2</sub>	3-pyridyl	phenyl
40	NH	4-pyrimidyl	phenyl
41	NR <sup>1</sup>	4-pyridyl	phenyl
42	NH	p-aminomethylphenyl	phenyl
43	NHCH <sub>2</sub>	4-aminophenyl	phenyl

Table 2

Compound No.	L	Ar'	R <sup>3</sup>
44	NH	4-pyridyl	3-chlorophenyl
45	NH	Phenyl	4-pyridyl
46	NH		phenyl
48	NH	2-benzylamino-3-pyridyl	phenyl
49	NH	2-benzylamino-4-pyridyl	phenyl
50	NH	3-benzyloxyphenyl	phenyl
51	NH	4-pyridyl	3-aminophenyl
52	NH	4-pyridyl	4-pyridyl
53	NH	4-pyridyl	2-naphthyl
54		4-pyridyl	phenyl
55		Phenyl	phenyl
56		2-pyridyl	phenyl
61	NH	4-pyridyl	2-trifluoromethyl phenyl
62	NH	4-aminophenyl	phenyl
64	NH	3-methoxyphenyl	2-fluorophenyl
65	NH	4-methoxyphenyl	2-fluorophenyl
66	NH	4-pyrimidinyl	2-fluorophenyl
67	NH	3-amino-4-pyridyl	phenyl
68	NH	4-pyridyl	2-benzylaminophenyl
69	NH	2-benzylaminophenyl	phenyl
70	NH	2-benzylaminophenyl	4-cyanophenyl
71	NH	3'-cyano-2-benzylaminophenyl	phenyl

A7  
T0300T = 2925242660

(b) the compounds listed in Table 3, below, wherein L is NH; Z<sup>3</sup> is N; Z<sup>6</sup> and Z<sup>7</sup> are CH and Z<sup>5</sup>, Z<sup>8</sup>, Ar' and R<sup>3</sup> are as shown in Table 3:

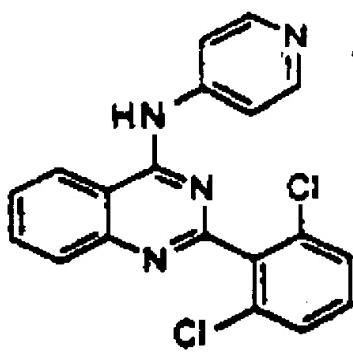
Table 3				
Compound No.	Z <sup>5</sup>	Z <sup>8</sup>	Ar'	R <sup>3</sup>
72	CH	N	4-pyridyl	2-fluorophenyl
73	CH	N	4-pyridyl	2-chlorophenyl
74	CH	N	4-pyridyl	phenyl
75	N	N	4-pyridyl	phenyl
76	N	CH	4-pyridyl	phenyl

and

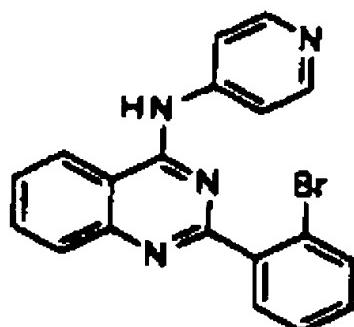
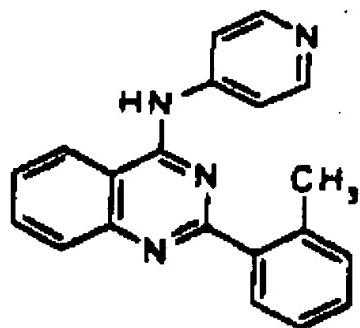
(c) the quinazoline derivatives listed in Table 4 below, wherein L is NH; Ar' is 4-pyridyl; Z<sup>3</sup>, Z<sup>5</sup>, and Z<sup>8</sup> are N; Z<sup>6</sup> or Z<sup>7</sup> are CR<sup>2</sup> as shown and each is otherwise N and wherein R<sup>3</sup> and R<sup>2</sup> are as shown in Table 4:

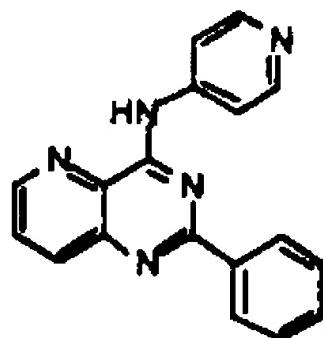
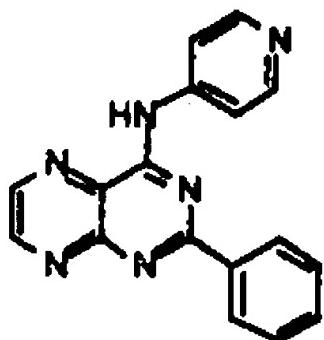
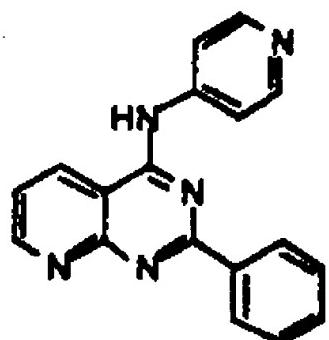
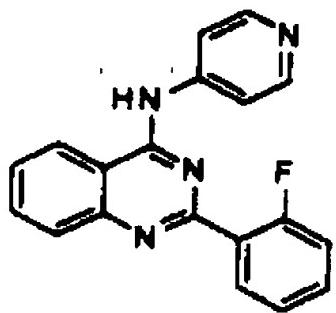
Table 4		
Compound No.	R <sup>3</sup>	R <sup>2</sup>
77	2-chlorophenyl	6,7-dimethoxy
78	2-fluorophenyl	6-nitro
79	2-fluorophenyl	6-amino
80	2-fluorophenyl	7-amino
81	2-fluorophenyl	6-(3-methoxybenzylamino)
82	2-fluorophenyl	6-(4-methoxybenzylamino)
83	2-fluorophenyl	6-(2-isobutylamino)
84	2-fluorophenyl	6-(4-methylmercaptophenylamino)
85	2-fluorophenyl	6-(4-methoxybenzoyl amino)
86	4-fluorophenyl	7-amino
87	4-fluorophenyl	7-(3-methoxybenzylamino)

31. (New) The composition of claim 18 wherein the compound of formula (1) is selected from the group consisting of the following compounds:



A7

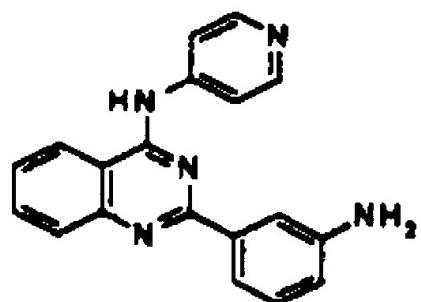
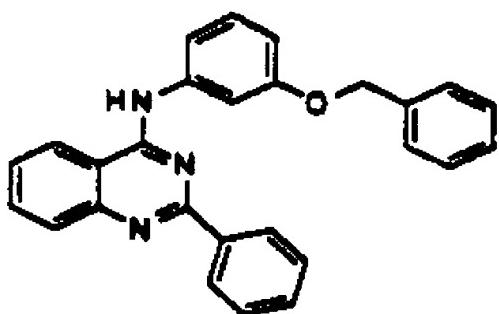
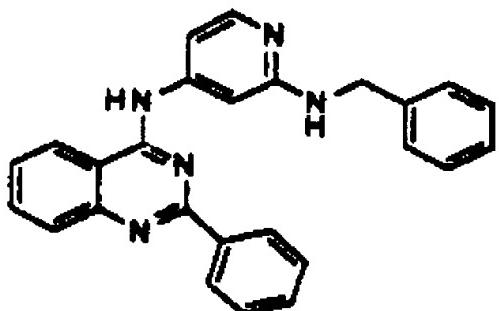
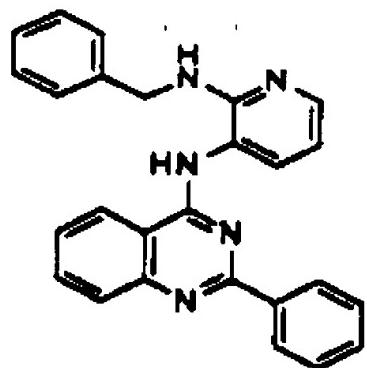


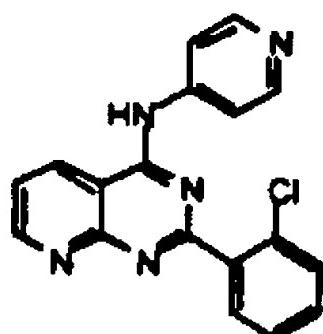
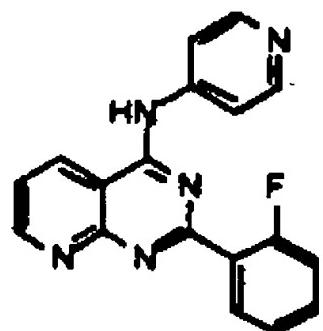
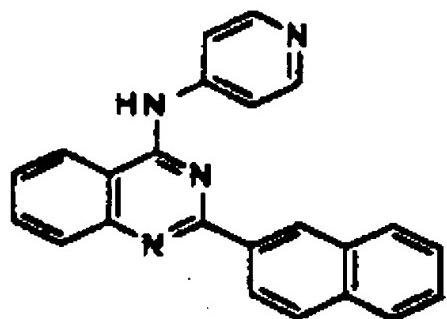
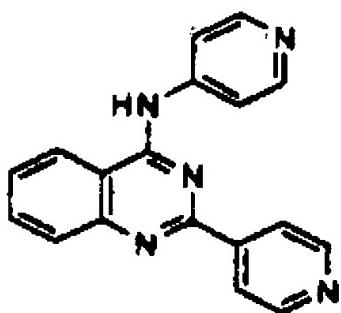


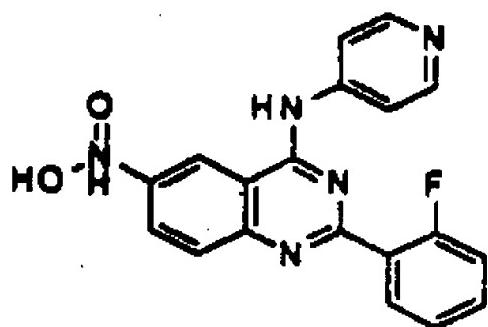
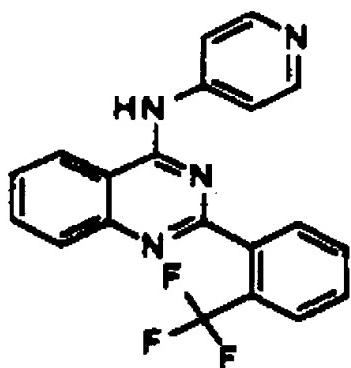
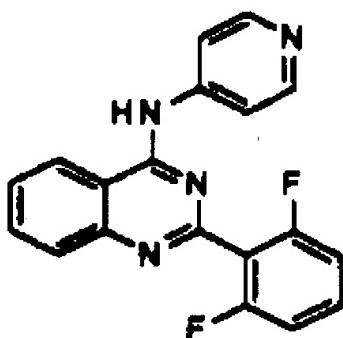
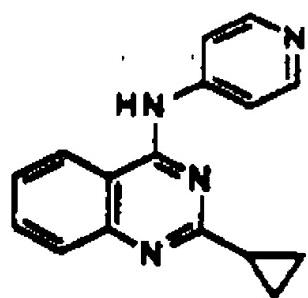
A7

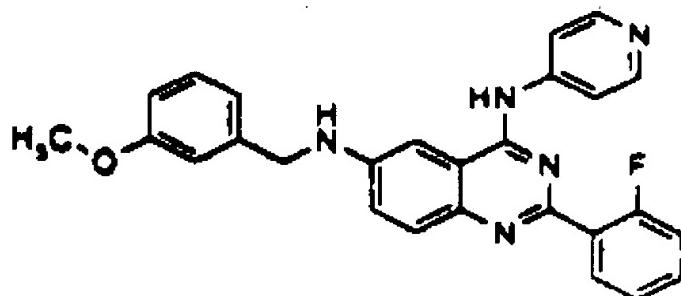
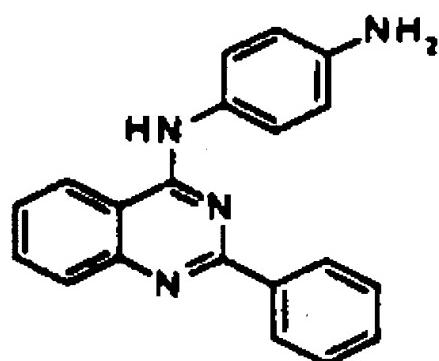
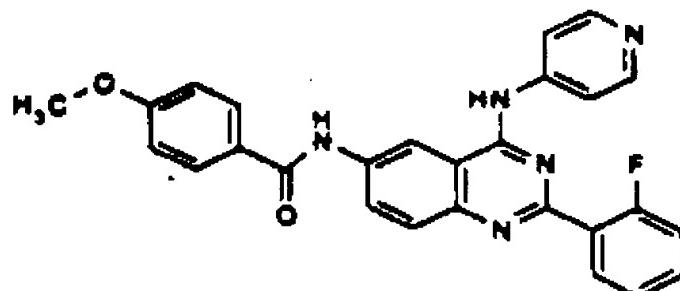
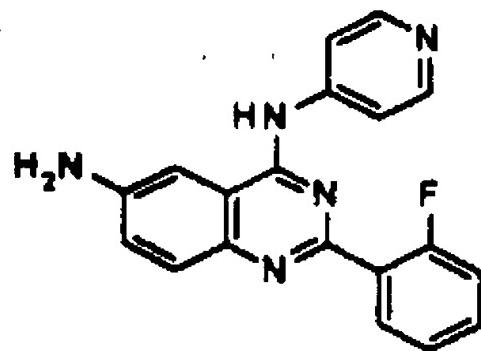
F0500TF-2352X560

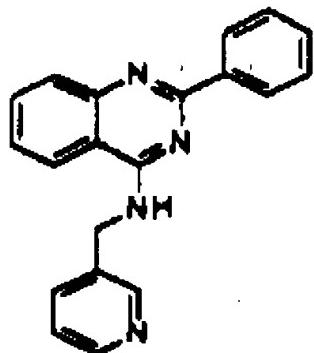
A7



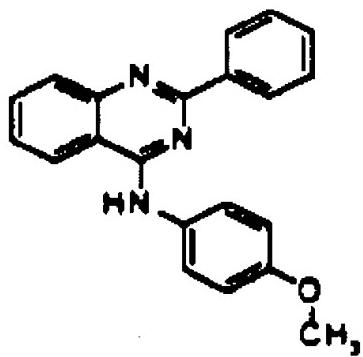
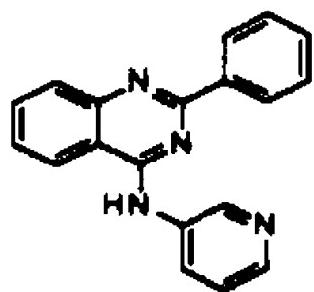
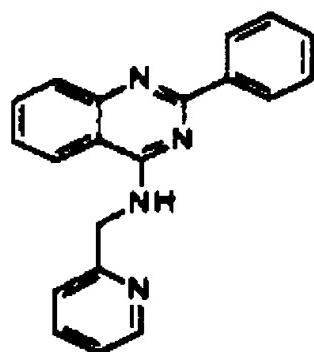


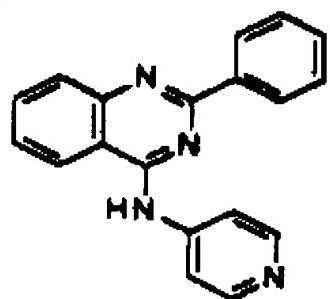
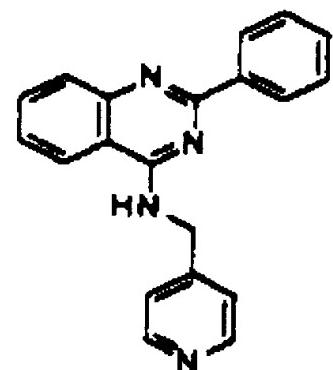
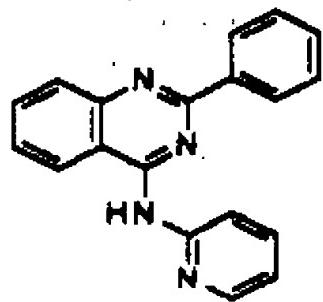




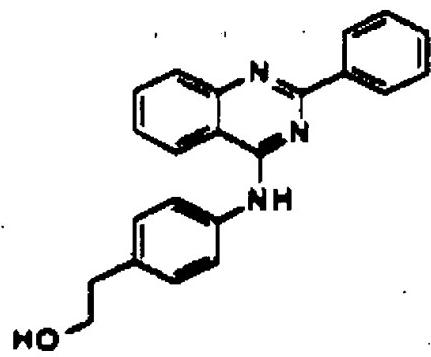


A7

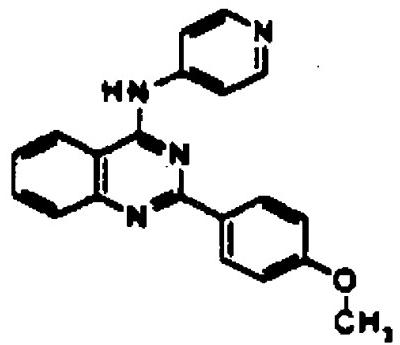
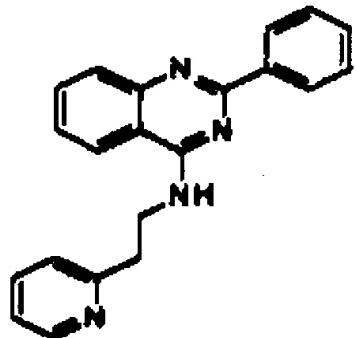
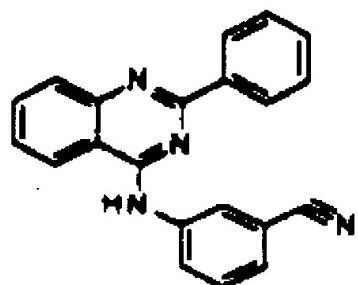




A7

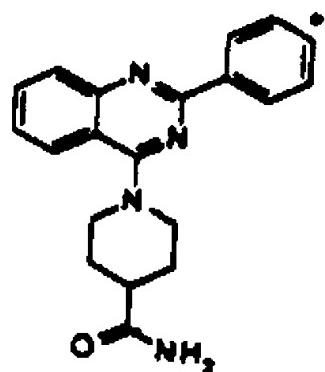
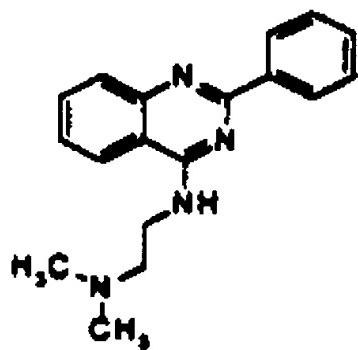
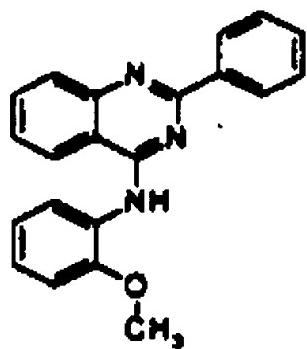


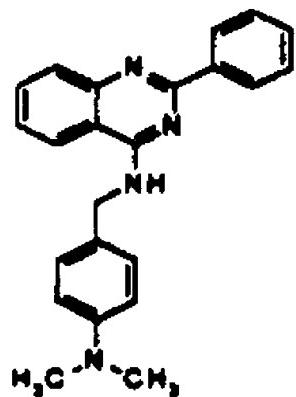
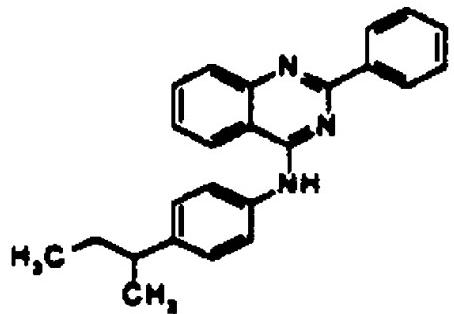
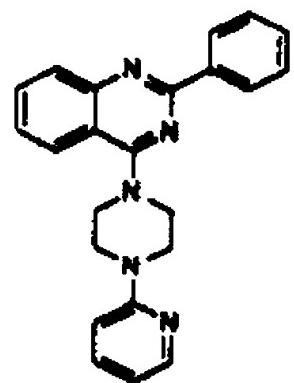
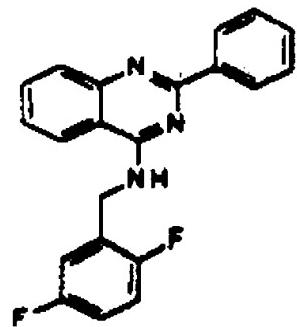
A7

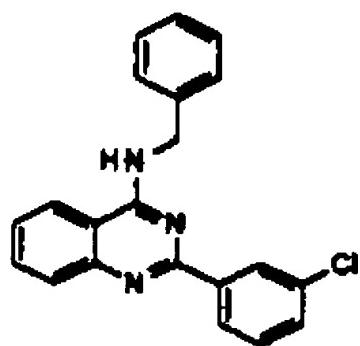


06972582-100501

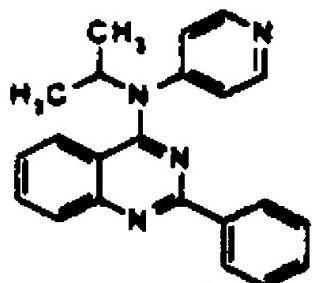
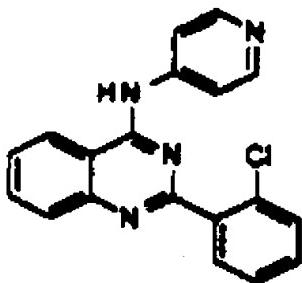
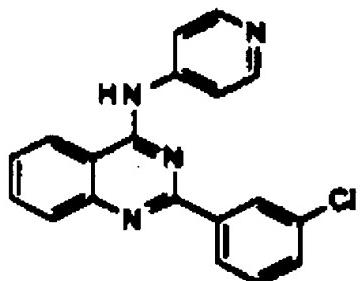
A7

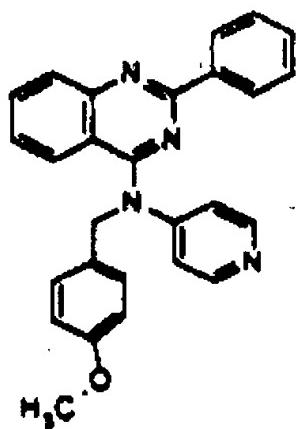




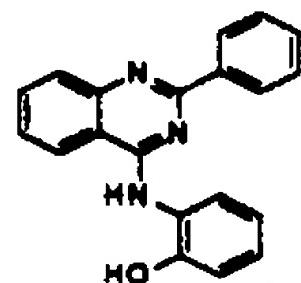
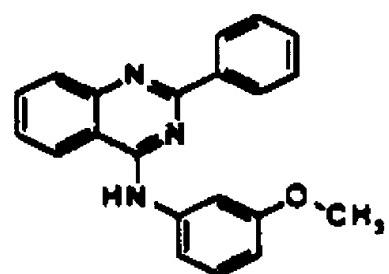
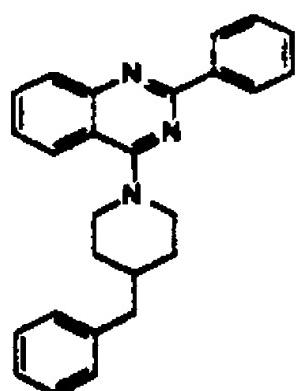


A7

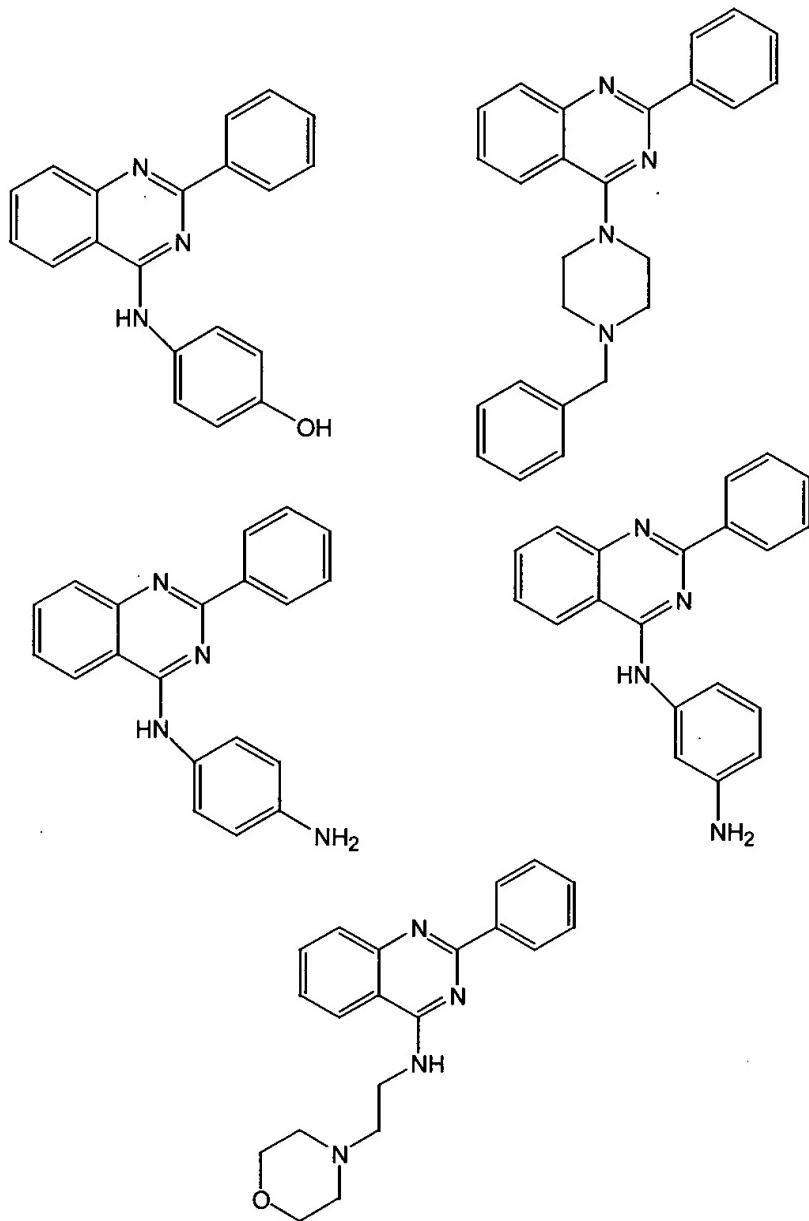




117

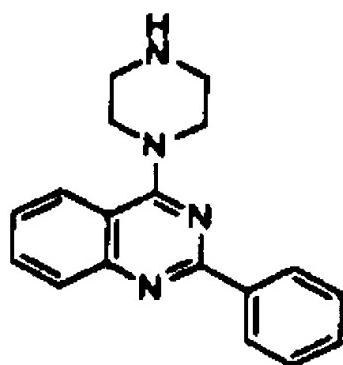
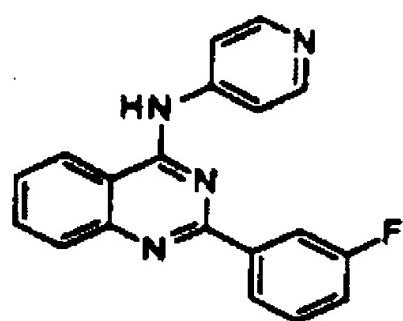
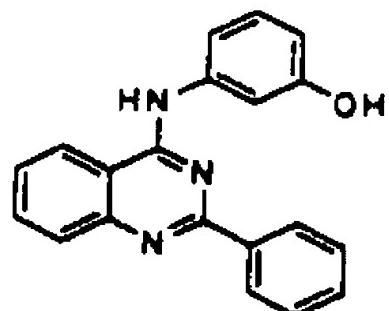


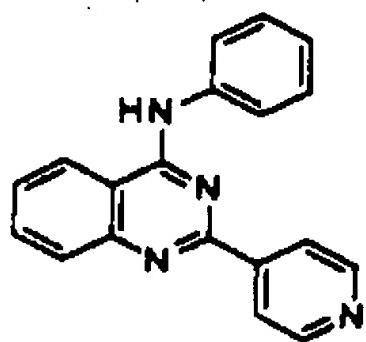
A7



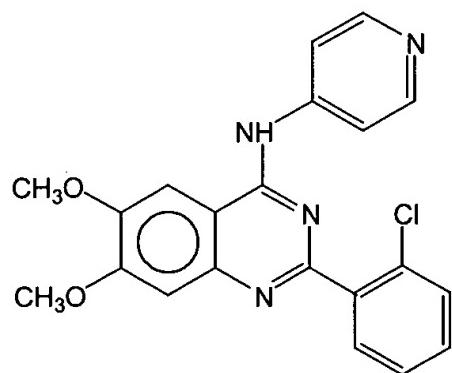
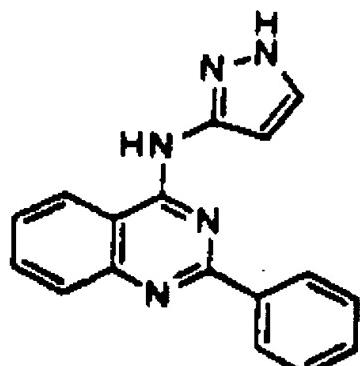
09522582-A00501

A7

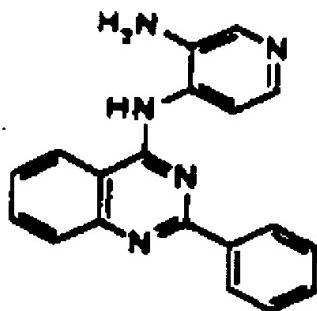
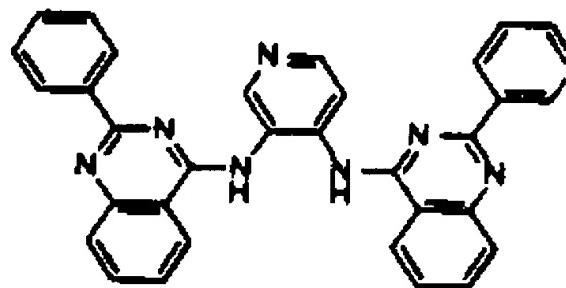
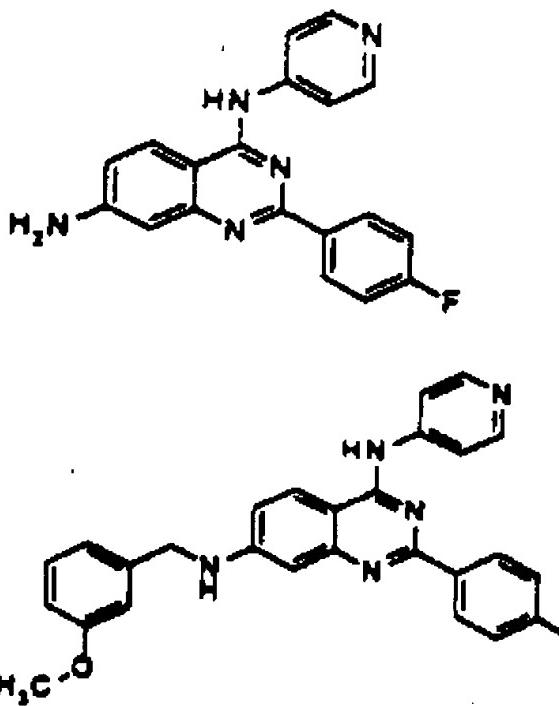




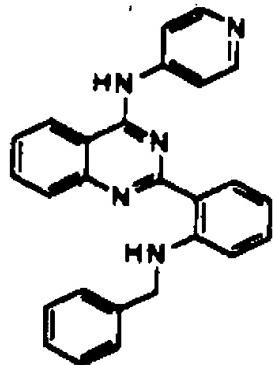
A1



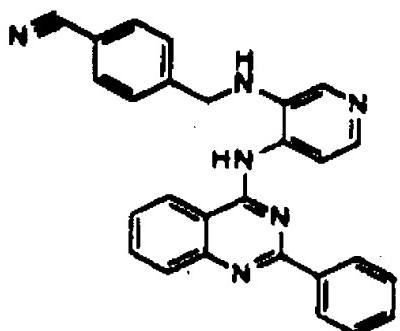
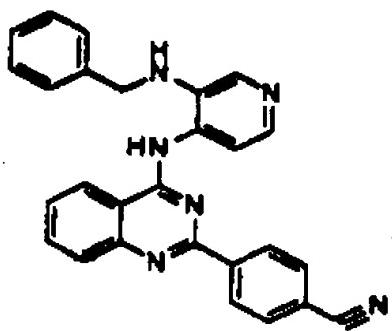
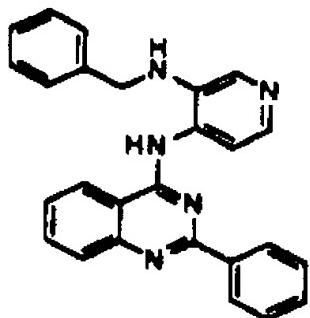
A7

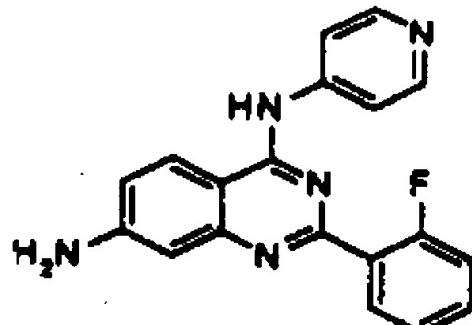
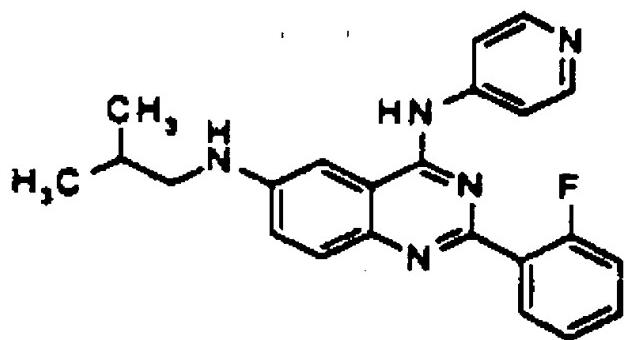


1005007-28527660

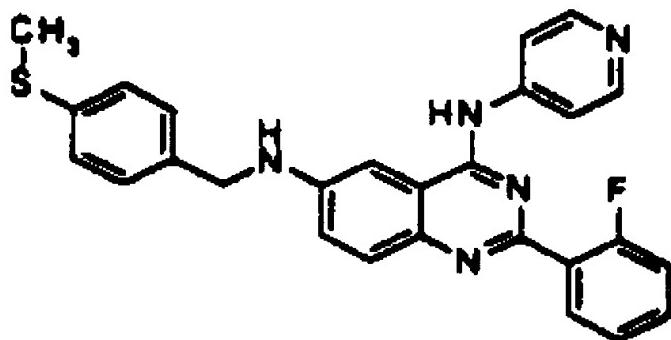
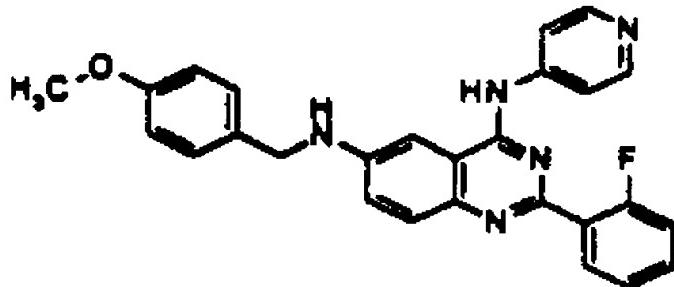


A7



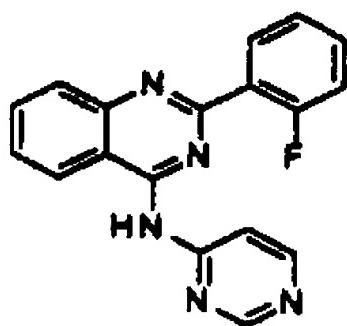
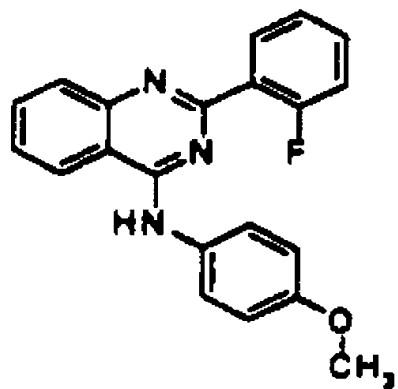
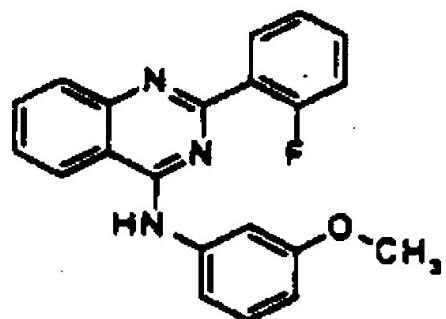


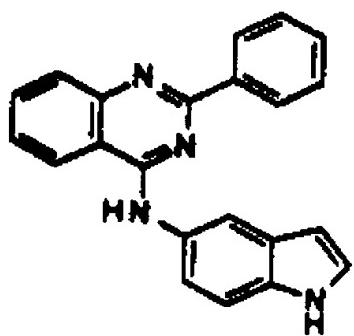
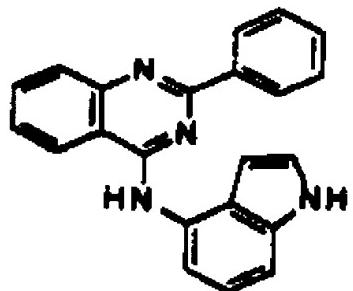
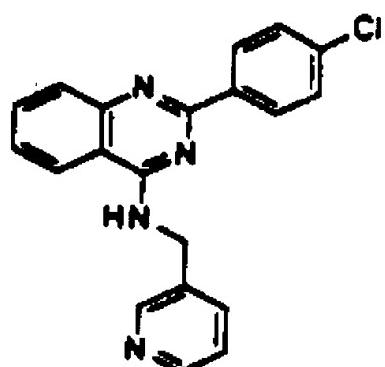
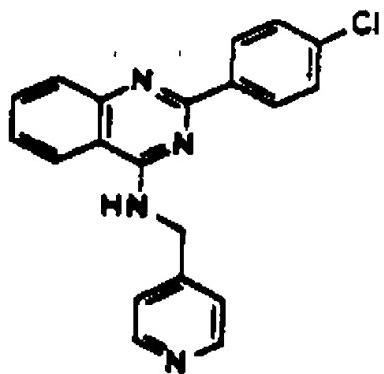
A7

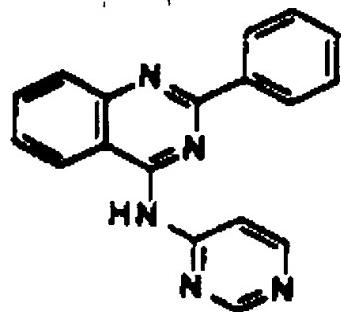


09972582 • 100501

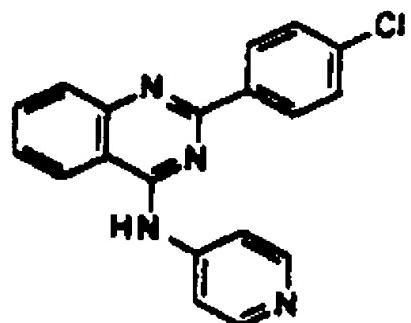
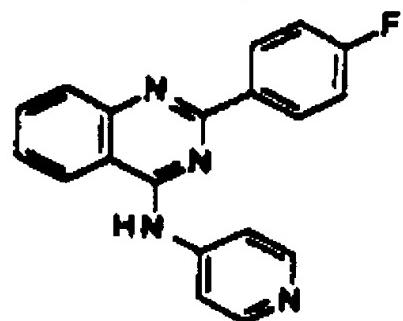
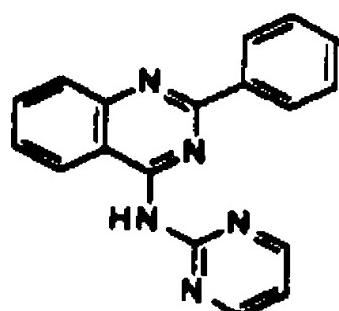
A7







A7



32. (New) The composition of claim 18 wherein

L is -R<sup>1</sup>N(CH<sub>2</sub>)<sub>n</sub>-;

L is -R<sup>1</sup>N(CH<sub>2</sub>)<sub>n</sub>- wherein R<sup>1</sup> is H or is alkyl (1-6C) or arylalkyl optionally substituted on the aryl group with 1-3 substituents independently selected from alkyl (1-6C), halo, OR, NR<sub>2</sub>, SR, -OOCR, -NROCR, RCO, -COOR, -CONR<sub>2</sub>, -SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or lower alkyl (1-4C) and n is 0, 1 or 2; and

(a) Ar' is phenyl, substituted with at least one group selected from the group consisting of optionally substituted alkyl (1-6C), halo, OR, NR<sub>2</sub>, SR, -OOCR, -NROCR, RCO, -COOR, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or lower alkyl (1-4C), or pyridyl, indolyl, or pyrimidyl, each optionally substituted with at least one group selected from the group consisting of optionally substituted alkyl (1-6C), halo, OR, NR<sub>2</sub>, SR, -OOCR, -NROCR, RCO, -COOR, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or lower alkyl (1-4C); and

R<sup>3</sup> is phenyl optionally substituted with 1-3 substituents which substituents are selected from the group consisting of alkyl (1-6C), halo, OR, NR<sub>2</sub>, SR, -OOCR, -NROCR, RCO, -COOR, -CONR<sub>2</sub>, -SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or lower alkyl (1-4C); or

(b) Ar' is phenyl, pyridyl, indolyl, or pyrimidyl, each optionally substituted with a group selected from the group consisting of optionally substituted alkyl (1-6C), halo, OR, NR<sub>2</sub>, SR, -OOCR, -NROCR, RCO, -COOR, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or lower alkyl (1-4C); and

R<sup>3</sup> is phenyl substituted with 1-3 substituents which substituents are selected from the group consisting of alkyl (1-6C), halo, SR, -OOCR, -NROCR, RCO, -COOR, -CONR<sub>2</sub>, -SO<sub>2</sub>NR<sub>2</sub>, CN, and CF<sub>3</sub>, wherein each R is independently H or lower alkyl (1-4C); or

(c) Ar' is phenyl substituted with a group selected from the group consisting of optionally substituted NR<sub>2</sub>, SR, -NROCR, RCO, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, and CF<sub>3</sub>, wherein each R is independently H or lower alkyl (1-4C); or pyridyl substituted with a group selected from the group consisting of optionally substituted alkyl (1-6C), halo, OR, NR<sub>2</sub>, SR, -OOCR, -NROCR, RCO, -COOR, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or lower alkyl (1-4C); or indolyl or pyrimidyl, each optionally substituted with a group selected from the group consisting of optionally substituted alkyl (1-6C), halo, OR, NR<sub>2</sub>, SR, -OOCR,

-NROCR, RCO, -COOR, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or lower alkyl (1-4C); and

R<sup>3</sup> is phenyl optionally substituted with 1-3 substituents which substituents are selected from the group consisting of alkyl (1-6C), halo, OR, NR<sub>2</sub>, SR, -OOCR, -NROCR, RCO, -COOR, -CONR<sub>2</sub>, -SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or lower alkyl (1-4C); or

(d) Ar' is phenyl, pyridyl, indolyl, or pyrimidyl, each optionally substituted with a group selected from the group consisting of optionally substituted alkyl (1-6C), halo, OR, NR<sub>2</sub>, SR, -OOCR, -NROCR, RCO, -COOR, -CONR<sub>2</sub>, SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or lower alkyl (1-4C); and

R<sup>3</sup> is phenyl substituted with 1-3 substituents which substituents are selected from the group consisting of alkyl (1-6C), halo, OR, SR, -OOCR, -NROCR, RCO, -COOR, -CONR<sub>2</sub>, -SO<sub>2</sub>NR<sub>2</sub>, CN, CF<sub>3</sub>, and NO<sub>2</sub>, wherein each R is independently H or lower alkyl (1-4C).

33. (New) The composition of claim 18 wherein the compound of formula 1 is selected from the group consisting of

2-phenyl-4-(4-pyridylamino)-quinazoline;  
2-(2-bromophenyl)-4-(4-pyridylamino)-quinazoline;  
2-(2-chlorophenyl)-4-(4-pyridylamino)-quinazoline;  
2-(2-fluorophenyl)-4-(4-pyridylamino)-quinazoline;  
2-(2-methylphenyl)-4-(4-pyridylamino)-quinazoline;  
2-(4-fluorophenyl)-4-(4-pyridylamino)-quinazoline;  
2-(3-methoxyanilyl)-4-(4-pyridylamino)-quinazoline;  
2-(2,6-dichlorophenyl)-4-(4-pyridylamino)-quinazoline;  
2-(2,6-dibromophenyl)-4-(4-pyridylamino)-quinazoline;  
2-(2,6-difluorophenyl)-4-(4-pyridylamino)-quinazoline;  
2-(2-fluorophenyl)-4-(4-pyridylamino)-6, 7-dimethoxyquinazoline;  
2-(4-fluorophenyl)-4-(4-pyridylamino)-6, 7-dimethoxyquinazoline;  
2-(2-fluorophenyl)-4-(4-pyridylamino)-6-nitroquinazoline;  
2-(2-fluorophenyl)-4-(4-pyridylamino)-6-aminoquinazoline;  
2-(2-fluorophenyl)-4-(4-pyridylamino)-7-aminoquinazoline;

A7

2-(2-fluorophenyl)-4-(4-pyridylamino)-6-(3-methoxybenzylamino)-quinazoline;  
2-(2-fluorophenyl)-4-(4-pyridylamino)-6-(4-methoxybenzylamino)-quinazoline;  
2-(2-fluorophenyl)-4-(4-pyridylamino)-6-(2-isobutylamino)-quinazoline; and  
2-(2-fluorophenyl)-4-(4-pyridylamino)-6-(4-methylmercaptophenylamino)-quinazoline.

1000000 - 2000000000